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TITLE: NRC/AMRMC Resident Research Associateship Program

PRINCIPAL INVESTIGATOR: Howard R. Gamble, Ph.D.

CONTRACTING ORGANIZATION: National Academy of Sciences

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Advisers to the Nation on Science, Engineering, and Medicine

# National Research Council RESEARCH ASSOCIATESHIP PROGRAM with U.S. Army Medical Research & Materiel Command

## **Annual Contract Technical Report**

Contract No. W81XWH-12-2-0018 Contract Period: 02/06/2012-02/05/2017 Report Period: 02/06/2012-02/05/2013 During the reporting period, the NRC conducted the following activities in support of the subject contract:

#### **Outreach and Promotion**

The promotional schedule to advertise the National Research Council (NRC) Research Associateship Programs included the following: 1) attendance at meetings of major scientific and engineering professional societies; 2) advertising in programs and career centers for these and other professional society meetings; 3) direct mailing and emailing of announcements and program materials to presidents, graduate deans, and heads of appropriate science and engineering departments and minority-affairs offices of all academic degree-granting institutions in the United States; 4) posting announcements on internet job sites, electronic newsletters and professional society websites; 5) print advertising in high profile publications (e.g., Science magazine, the Chronicle of Higher Education); and, 6) maintaining a presence on social media sites such as Facebook.

The NRC attended a number of minority focused events in which we maintained exhibit booths, participated in workshops and advertised in meeting literature, newsletters and websites or submitted materials for distribution. In addition, ads were placed in a variety of minority publications (e.g., Affirmative Action, Black Collegian).

In advertising the Research Opportunities available to prospective applicants, the NRC maintained an up-to-date listing of all active Research Advisers, current Adviser contact information and details of each Research Opportunity.

#### **Processing and Review of Applications**

Applications to the Research Associateship Program were submitted via a web-based application system. Each of the four application cycles opened two months prior to the application deadline. NRC staff provided support to prospective applicants including providing application instructions, technical support and additional information as requested.

A summary of applications for the reporting period is shown in Table 1.

For each applicant, the NRC received and processed an application form, a research proposal, transcripts, a statement of previous and current research, and confidential reference reports. An application file check was made prior to the review and each applicant was notified if required documents were missing.

The NRC convened panels in five broad discipline areas for the competitive review of applications in the Research Associateship Programs. Results of the review were made available to Laboratory Program Representatives immediately following the conclusion of the each review.

A summary of the outcome of the review of applications for the reporting period is shown in Table 1.

#### **Administration of Awards**

The NRC made awards to applicants based on sponsor authorization. A summary of awards authorized and the acceptance or declination by the applicant during the current reporting period is shown in Table 1.

For Associates beginning or continuing tenure, the NRC provided the administrative functions described in the contract Statement of Work. These functions included stipend payments, management of a major medical benefits insurance program, and reimbursement for relocation and travel to professional meetings.

A summary of NRC Research Associates on tenure during the reporting period is shown in Table 2.

#### **Outcomes Reporting**

All NRC Associates who completed tenure were required to submit a final report that described the outcome of their Associateship award. Final reports received by the NRC during the current reporting period are attached to this technical report.

The activities of Associates submitting final reports during this reporting period, including publications, presentations and patents, as well as an assessment of their experience in the program, are summarized in Table 3. Specific research accomplishments of Associates completing tenure during the reporting period are summarized in Table 4.

**Table 1.** Applications and Awards

**Table 2.** Associates on Tenure

**Table 3.** Associates Activity

**Table 4.** Summary of Associate Research

**Attachments:** Associate Final Reports

## U.S. Army Medical Research & Materiel Command <a href="Table 1: Applications and Awards">Table 1: Applications and Awards</a>

	May 2012	Aug 2012	Nov 2012	Feb 2013	Total
TOTAL APPLICATIONS	8	15	5	8	36
Applications not reviewed	1	1	2	2	6
Applications reviewed	7	14	3	6	30
Not recommended	0	0	0	0	0
Recommended	7	14	3	6	30
Withdrawn	0	0	0	0	0
Lab decision pending	0	0	2	6	8
Awards offered	7	9	1	0	17
Applicant decision pending	0	0	0	0	0
Awards accepted	7	9	1	0	17
Awards declined	0	0	0	0	0
Not funded	0	5	0	0	5

## **Table 2: Associates on Tenure**

Associate	Adviser	Tenure Dates	Country of Citizenship	Final Report
U.S. Army Institute of Surgical Resea				
Burdette, Alexander Justin	Alvarez, Rene	10/1/2012-9/30/2013	United States	
Chen, Guojun	Devore, David I	8/8/2011-5/14/2012	China	Received
Cheppudira, Bopaiah Pooviah	Clifford, John L	9/4/2012-9/3/2013	India	
Childers, Brandon Michael	Leung, Kai P	7/11/2011-12/31/2012	United States	Not Recv'd
Choi, Jae hyek	Wang, Heuy-Ching H.	11/15/2011-4/14/2013	Korea, South	
Flagg, Shannon Coates	Alvarez, Rene	5/14/2012-5/13/2013	United States	
Greene, Whitney Ann	Wang, Heuy-Ching H.	4/25/2012-4/24/2014	United States	
Hurtgen, Brady Joe	Wenke, Joseph C	12/3/2012-12/2/2013	United States	
Kaini, Ramesh Raj	Wang, Heuy-Ching H.	1/3/2013-1/2/2014	United States	
Kreyer, Stefan Franz Xaver	Batchinsky, Andriy I	12/7/2012-12/6/2013	Germany	
Langer, Thomas	Cancio, Leopoldo C.	8/29/2011-11/28/2012	Italy	Received
Meledeo, Michael Adam	Bowman, Phillip Dalton	3/1/2010-7/6/2012	United States	Received
Miller, Christine Lindsay	Leung, Kai P	2/4/2013-2/3/2014	United States	
Muniz, Alberto	Wang, Heuy-Ching H.	10/14/2008-10/13/2013	United States	
Nyland, Jennifer Elaine	Clifford, John L	2/1/2013-1/31/2014	United States	
Parida, Bijaya Kumar	McFaul, Steve J.	3/19/2012-3/18/2014	India	
Pilia, Marcello	Rathbone, Christopher R	9/4/2012-9/3/2013	United States	
Rose, Lloyd Frederick	Leung, Kai P	11/1/2012-10/31/2013	United States	
Salas, Margaux Marie	Clifford, John L	10/10/2012-10/9/2013	United States	
Samberg, Meghan	Christy, Robert John	10/1/2012-9/30/2013	United States	
Sanchez, Carlos Jose	Wenke, Joseph C	9/7/2011-9/6/2013	United States	
Scaravilli, Vittorio	Batchinsky, Andriy I	11/15/2012-11/14/2013	Italy	
Schneider Herrera, Bruno	Leung, Kai P	5/1/2012-4/30/2013	Brazil	
Torres, Luciana Neves	Dubick, Michael A.	5/18/2011-8/17/2012	Brazil	Received
Van Laar, Tricia Annette	Leung, Kai P	4/9/2012-4/8/2014	United States	110001100
Vecchi, Vittoria	Cancio, Leopoldo C.	11/14/2011-10/22/2012	Italy	Received
Ward, Catherine Lindsey	Wenke, Joseph C	8/22/2011-8/21/2013	United States	received
U.S. Army Medical Research Institute		0/22/2011-0/21/2013	Officed States	
Hubbard, Kyle	McNutt, Patrick Michael	6/1/2012-5/31/2013	United States	
Wong, Benjamin Jay	Sciuto, Alfred Mario	2/7/2011-9/4/2012	United States	Received
Yego, E Chepchumba Koech	Dillman, James F.	9/1/2010-6/30/2013	United States	received
U.S. Army Medical Research Institute		9/1/2010-0/30/2013	Officed States	
Bounds, Callie Elizabeth	Schmaljohn, Connie	3/1/2011-2/28/2014	United States	
Brocato, Rebecca Lee	Hooper, Jay W.	1/19/2011-3/13/2013	United States	
Copeland, Anna Maria	Schmaljohn, Connie	1/18/2011-1/17/2014	United States	
Heffron, Jared David	Welkos, Susan Lee	8/16/2010-3/15/2013	United States	
Johnston, Sara Christine	Goff, Arthur James	3/25/2009-5/1/2012	United States	Received
Kwilas. Steven Alexander	· · · · · · · · · · · · · · · · · · ·	4/13/2010-4/12/2014	United States	received
,	Hooper, Jay W.	3/1/2011-2/28/2014		
Lindquist, Michael Edward	Schmaljohn, Connie		United States	
Maes, Piet	Hooper, Jay W.	12/3/2012-12/2/2013	Belgium	
Morazzani, Elaine Marie	Glass, Pamela J	12/1/2011-11/30/2014	United States	
Rahman, Md. Mizanur	Ahmed, Syed Ashraf	3/1/2011-2/28/2014	Bangladesh	
Trefry, John Christopher Walter Reed Army Institute of Resear	Honko, Anna Nichole	7/1/2011-6/30/2013	United States	
Abente, Eugenio	Bodhidatta, Ladaporn	9/17/2012-9/16/2013	United States	
Boutte, Angela	Dave, Jitendra R.	6/15/2011-1/14/2013	United States	Received
Caudle, Krista Layne	Tortella, Frank Casper	1/28/2013-1/27/2014	United States	1/CCGIVEU
			United States United States	
Kobylinski, Kevin Conrad	Romero, Gabriela Zollner	10/17/2011-10/16/2013		Received
Leung, Lai Yee	Tortella, Frank Casper	2/16/2010-1/31/2013	Hong Kong	Received
Linton, Yvonne-Marie	Wilkerson, Richard C.	10/3/2011-10/2/2013	United Kingdom	Dozahirat
McCoy, Margaret Ellen	Lanar, David Ervin	7/27/2009-10/4/2012	United States	Received
Melendrez, Melanie Crystal	Jarman, Richard George	3/22/2010-5/21/2012	United States	Received
Moore, Nicole LT	Genovese, Raymond F.	7/29/2010-7/28/2013	United States	
Mountney, Andrea	Tortella, Frank Casper	11/15/2010-2/28/2013	United States	
Mutihac, Radu	Balkin, Thomas J.	3/1/2011-2/28/2013	Romania	
Pichard, Luis Eduardo	Wesensten, Nancy J.	7/24/2012-7/23/2013	United States	

Contract Technical Report Contract # W81XWH-12-2-0010

Report Period: 02/06/2012-02/05/2013

U.S. Army Medical Research & Materiel Command 3/18/2013, 3:21 PM

Pichugin, Alexander Vladimirovich	Krzych, Urszula	1/12/2009-7/11/2012	Russia	Received
Rajendran, Gnana Ravi	Kozar, Michael Patrick	9/7/2010-8/31/2012	United States	Received
Rossetti, Franco	Yourick, Debra Lynn	6/1/2009-5/31/2013	Brazil	
Sambanthamoorthy, Karthik	Luo, Chunyuan	3/1/2011-2/28/2014	India	
Yoon, K. Lira	Balkin, Thomas J.	1/3/2013-1/2/2014	United States	
Zarling, Stasya Nicole	Krzych, Urszula	2/7/2011-2/6/2014	United States	

#### **Table 3: Associates' Activities**

- 14 Associates ended tenure during the report period
- 24 months was the average tenure length
- **42** months was the longest
- 9 months was the shortest
- 13 submitted final reports

In the final reports, Associates indicated the following scholarly activity while on tenure.

- 17 Articles published in refereed journals
- **0** Patent applications
- 6 International presentations
- 34 Domestic presentations
- 3 Awards

After ending their tenure, Associates indicated their future plans as follows:

- 1 Permanent position at the NRC host agency
- **6** Contract or temporary position at the NRC host agency
- 1 Research/administrative position with another U.S. government agency
- 1 Research/administrative position with foreign government agency
- 1 Research/teaching at US college/university
- 1 Research/teaching position at a foreign college or university
- **0** Research/administrative position in private industry in the U.S.
- **0** Research/administrative position in private industry outside of the U.S.
- **0** Research/administrative position with a non-profit
- 0 Self-employed/consulting
- 2 Postdoctoral Research
- 0 Other
- **0** No information provided

In their final reports, Associates were asked to evaluate certain aspects of their experiences on a scale of 1 (low) to 10 (high). The average rating for each item follows:

- 9.4 Short-term value (lab)-Development of knowledge, skills, and research productivity at lab
- 9.2 Long-term value (career)-How your Research Associateship affected your career to date
- **8.8** Laboratory Support-Equipment, funding, orientation, safety & health training, etc.
- 9.5 Adviser Mentoring-Quality of mentoring from the Research Adviser
- 9.2 LPR Support-Quality of administrative support from the LPR
- 9.2 NRC Support-Quality of administrative support from the NRC

## **Table 4: Summary of Associate Research**

	Associate	Tenure Dates					
Bou	tte, Angela	6/15/2011-1/14/2013					
1	Determined the time course of brain der penetrating ballistic-like brain injury (PB		in brain tissue and serum of a rat model of				
2	Determined the time course of brain der traumatic stress disorder (fear condition		in brain tissue and serum of a rat model post- ping)				
3	Determined the effect of therapeutic dru	gs to prevent increases in	brain and serum biomarkers after PBBI				
4	Initiated proteomics analysis of a rat mo	del post-traumatic stress o	disorder (PTSD) and blast injury (on going)				
5	(Combat Casualty Care Program), co-authored pilot grant applications to determine efficacy of therapeutics in PBBI.						
6	Publications - Under the guidance of Dr. Dave, our lab published our proteomics study in the journal Electrophoresis. In a separate cohort using proteomics and bioinformatics prediction of biomarkers, I aided in defining novel biomarkers of PBBI in brain tissue that I have independently confirmed. A bioinformatics-themed manuscript is complete and pending submission.						
Che	n, Guojun	8/8/2011-5/14/2012					
1	Virus-like particle for antisense ODN de	livery					
2	Peptide-oligonucleic acid conjugates for						
3	Co-polymer enhanced liposome for anti						
Joh	nston, Šara	3/25/2009-5/1/2012					
1	MPXV Active Disease Surveillance Prog	gram identifying significant	increases in prevalence in DRC				
2	Identification of potential novel therapeu						
3	Identification of MPXV variants actively						
4	In vitro and in vivo evaluation of novel the		aviruses				
5	Development and implementation of dia	gnostic ELISA assays aga	ainst MPXV				
Lan	ger, Thomas	8/29/2011-11/28/2012					
1	Development of a large animal model to	study spontaneous breatl	hing during extracorporeal gas exchange				
2	y Distress Syndrome	•	s and animals with the Acute Respirator				
3	·						
4			esults in healthy in the Acute Respiratory Distress and monitoring of pulmonary disease reducing				
Leu	ng, Lai Yee	2/16/2010-1/31/2013					
1	future neuroprotective drug studies.		emorrhagic hypotension that will be used for				
2	found to be unique under different injury	combinations.	a models. The patterns of these changes were				
3	following PBBI whereas hypoxemia only	prolonged the depolariza					
4							
5	The sequence of PBBI, hemorrhagic hypotension and hypoxemia affected the mortality rate and neurological deficits. PBBI, HS followed by HX resulted in the highest mortality rate (50%) and more neurological deficits among all polytrauma groups.						
	Coy, Margaret	7/27/2009-10/4/2012					
1	Identified the mechanism of action of SA						
2	Designed and carried out experiments t within the immune system	o examine and characteriz	te the processing and presentation of SAPN				

- 3 Examined the phenotypes of SAPN-specific T-cell populations and proved that CD8+ T-cells from mice immunized with SAPN are able to, by themselves, induce sterile immunity in mice- this is the first malaria vaccine to be able to show this.
- 4 Examined needleless approcahes to vaccine delivery, including pleuronic lecithin organogel creams
- 5 Identified potential effects on vaccine efficacy resulting from the addition of mosquito saliva
- 6 The youngest investigator at the WRAIR to successfully write and complete a non-human primate trial for malaria

#### Meledeo, Michael

#### 3/1/2010-7/6/2012

- Aptamers can be used to completely inhibit the anti-coagulant effects of activated protein C (aPC); however, aPC does not appear to be the sole sufficient cause of the acute coagulopathy of trauma.
- 2 Exposing in vitro cultures of endothelial cells (ECs) to laminar flow (as a model of their physiological environment) will induce a number of changes to both EC morphology and gene expression in a variety of inflammatory and morphology pathways.
- While not all of the changes in gene expression result in an altered proteome, there are a number of significant differences in protein expression between static cultured ECs and those exposed to flow.
- 4 An analysis of the EC glycocalyx through confocal microscopy and western blotting of membrane proteins and associated glycoforms has led to an enhanced understanding of the structure and function of the endothelial glycocalyx layer.
- All of these have provided advancement in the formulation of in vitro models of endothelium; in the future it should be possible to use these models as a platform for the testing of both therapeutics and diagnostics for vascular dysfunction.

#### Melendrez, Melanie

#### 3/22/2010-5/21/2012

- 1 Variants analysis of dengue quasispecies populations showed that variants are host or vector specific despite containing the same consensus sequence. Diversity was found to not be constrained within the vector as suggested in some publications.
- Selection analysis showed the populations to be expanding, evolving at a faster rate than 'average' for the dataset, and were under selective pressure with a predominance of nonsynonymous mutations when compared with the consensus sequence.
- Phylogenetic analysis revealed that dengue quasispecies sequences isolated in 2010 were distinct from other ciruclating consensus sequences from Thailand and full E gene offered higher resolution than partial E gene sequences.
- 4 Amino acid (aa) analysis suggested several positions where changes would affect replication, antibody binding or VLP assembly according to the literature. Multivariate analysis predicted uncharacterized aa positions that would have impact if altered.
- This work revealed the importance of full E gene surveillance for assessment of an changes, illustrated the variability and pathogenic potential of dengue quasispecies variant diversity and established a baseline in which to make future comparisons.

#### Pichugin, Alexander

#### 1/12/2009-7/11/2012

- 1 10 novel liver stage Pb antigens reduce LS and BS parasite burden in C57Bl/6 mice.
- 2 3 novel liver stage Pb antigens sustain protection during 6 months after the last immunization.
- 3 | 3 novel liver stage Pb antigens enhance protection induced by PbCSP.
- 4 Established caged MHC-tetramer technology to use for discovery of T cell epitopes in malaria antigens.
- 5 Identified 3 immunodominant CD8 T cell epitopes from Pb PEVA.

#### Rajendran, Gnana

#### 9/7/2010-8/31/2012

- Several compounds were synthesized by the SAR of DQ and submitted for in vitro testing against blood stage malaria, specifically P.falciparum D6, W2, C235 and C2B strains, and assessed for metabolic stability in the mouse and human microsomes
- 2 In many cases the compounds solubility was improved but the compounds either lost potency against the C2B resistant strain of malaria or microsomal stability
- A few interesting trends were discovered. An unprecedented ester replacement, to the ethyl or morpholine amide was discovered that maintained potency against D6,W2,C235 but unfortunately lost activity against the key C2B atovaquone resistant strain
- 4 Additionally, many compounds were synthesized that maintained in vitro potency with significantly lower clogP (main focus of the research effort)
- 5 Efforts are currently toward acquiring a complete set of data to select profile compounds to be scaled up and tested in in vivo models

#### Torres, Luciana

#### 5/18/2011-8/17/2012

Intravital Microscopy was successfully employed for investigating EG shedding in hemorrhagic shock/resuscitation for the very first time;

Contract Technical Report Contract # W81XWH-12-2-0010 Report Period: 02/06/2012-02/05/2013 U.S. Army Medical Research & Materiel Command 3/18/2013, 3:21 PM

Intravital microscopy integrated with systemic hemodynamics evaluations may be essential and more accurate tools to identify changes and study mechanisms of EG shedding and systemic responses to hemorrhage and resuscitation therapy; Compared to baseline and to the sham group, there was a 50% reduction in endothelial glycocalyx (EG) thickness after hemorrhage and 60% increase in the levels of plasma Syndecan-1; Although resuscitation with LR and Hextend could stabilize hypotensive rats hemodynamicaly, these fluids were unable to restore EG thickness or coagulopathy (weak clots and prolonged coagulation time); Rats who received fresh frozen plasma (FFP) restored venular EG thickness to baseline level in addition to improve the systemic hemodynamics and coagulation response (restored homeostasis). Vecchi, Vittoria 11/14/2011-10/22/2012 Effects of radiation dose reduction on lung quantitative CT scan results in healthy in the Acute Respiratory Distress Syndrome: low-dose chest CT as a valuable tool for quantification and monitoring of pulmonary disease reducing patient exposure Use of quantitative CT for in vivo lung weight measurement: evaluate and monitor the time course of lung edema in ARDS measuring lung weight by gCT Pressure-guided positioning of bicaval dual-lumen catheters for veno-venous extracorporeal gas exchange Low-flow extracorporeal gal exchange for the treatment of ARDS caused by smoke inhalation and cutaneous burn 5 Extracorporeal Gas Exchange in awake spontaneously breathing sheep before and after the induction of ARDS Wong, Benjamin 2/7/2011-9/4/2012

1 Developed and characterized novel system for inhalational exposure of conscious animals to chemical agents

Utilized above system to examine the toxicokinetics (TK) of nerve agents and their analogs in rats
 Investigated utility of bronchodilators in a treatment regimen for inhalational chemical agent exposure

Advisers to the Nation on Science, Engineering, and Medicine National Research Council

## **Research Associateship Programs**

#### FINAL REPORT

1) Ass	sociate Last or Family Name		First Name		M.I.
Bout	te		Angela		M
3) To	day's Date		Dates of Tenure		
,	,				
10 JA	AN 2013		from 15 JUN 20	011 to 14 JAN 2013	
4)	Host Agency	Laboratory or Center		Division / Directorate / Departmen	nt
	AMRMC	WRAIR		Bran Trauma, Neuroprotection, N	leuro-
	(e.g., AFRL)	(e.g., Wright Patterson AFB)	)	restoration	
				(e.g., High-Speed Propulsion)	
5) <i>Na</i>	me of Laboratory NRC Advise	r (and USMA Mentor, if applicable)			
J	ITENDRA R. DAVE				

#### 6) TITLE OF RESEARCH PROPOSAL

#### **Biomarkers of Traumatic Brain Injuries**

#### 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.

- 1) Determined the time course of brain derived biomarkers elevated in brain tissue and serum of a rat model of penetrating ballistic-like brain injury (PBBI) (on-going)
- 2) Determined the time course of brain derived biomarkers elevated in brain tissue and serum of a rat model post-traumatic stress disorder (fear conditioning) and blast injury (on going)
- 3) Determined the effect of therapeutic drugs to prevent increases in brain and serum biomarkers after PBBI
- 4) Initiated proteomics analysis of a rat model post-traumatic stress disorder (PTSD) and blast injury (on going)
- 5) Grants Submitted grant proposal to define the markers and mechanism of severe and mild long term brain injury (Combat Casualty Care Program), co-authored pilot grant applications to determine efficacy of therapeutics in PBBI.
- 6) Publications Under the guidance of Dr. Dave, our lab published our proteomics study in the journal Electrophoresis. In a separate cohort using proteomics and bioinformatics prediction of biomarkers, I aided in defining novel biomarkers of PBBI in brain tissue that I have independently confirmed. A bioinformatics-themed manuscript is complete and pending submission.

## (USMA Davies Fellow: please add summary of teaching, including classes taught.) $\ensuremath{N\!/\!A}$

#### 8) RESEARCH IN PROGRESS Describe in no more than 100 words.

Post traumatic stress disorder (PTSD), traumatic brain injuries (TBIs) (e.g. blast overpressure (BOP), concussive, or penetrating) affect thousands military personnel. To successfully identify a panel of protein biomarkers that can (1) predict onset/existence and (2) define mechanisms, I am continuing studies in multiple brain regions and bio-fluids. Initial testing of preparative proteomics protocols was successful; each anatomical region (beginning with cerebral cortex) will be analyzed using proteomics and immunological methods. Biomarker abundance changes in tissues and biofluids after penetrating ballistic-like brain injury (PBBI) are being expanded to include therapeutic and sub-acute effects and we have fully developed studies to define biomarkers of mild, concussive, TBI.

#### 9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.

a) Publications in peer-reviewed journals

PROTEOMIC ANALYSIS AND BRAIN SPECIFIC SYSTEMS BIOLOGY IN A RODENT MODEL OF PENETRATING BALLISTIC-LIKE BRAIN INJURY Electrophoresis. 2012 Dec;33(24):3693-704.

Angela M Boutté, Changping Yao, Firas Kobaissy, Xi-Chun May Lu, Zhiqun Zhang, Kevin K.

Wang, Kara Schmid, Frank C. Tortella and Jitendra R. Dave

b) Books, book chapters, other publications
Protein Biomarkers in Traumatic Brain Injury: An Omics Approach Angela Boutte
c) Manuscripts in preparation, manuscripts submitted
1. SYSTEMS BIOLOGY META-ANALYSES OF GENOMIC DATASETS TO IDENTIFY CONSERVED MECHANISMS AND NOVEL BIOMARKERS OF TRAUMATIC BRAIN INJURY.
2. Time dependant biomarker abundance in brain tissue and serum after acute PBBI
3. Spatial-temporal biomarkers in a model of blast with and without fear conditioning stress in brain tissue and serum after acute and chronic injury
4. The effect of blast injury with and without fear conditioning on the cerebral cortex proteome
10) PATENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH Provide titles, inventors, and dates of applications.
N/A
11) PRESENTATIONS AT SCIENTIFIC MEETINGS OR CONFERENCES Provide complete references: author(s), title, abstract/proceeding citation, meeting name and location.
International
N/A
Domestic 1. Boutte A, et.al. (2012) Differential Protein Changes in Penetrating and Non-penetrating Models of Traumatic Brain Injury.  National Neurotrauma Society Symposium, Abstract #A48.
2. Yu C, Boutte A, et. al. (2012) Systems Biology and Meta-Analysis of Genomic Datasets to Identify Conserved Mechanisms and Novel Biomarkers of Traumatic Brain Injury. National Neurotrauma Society Symposium, Abstract # B48.
12) SEMINARS OR LECTURES DELIVERED AT UNIVERSITIES AND/OR INSTITUTES Include dates, names and locations of seminars.
1. Boutte A, et.al. (2012) Differential Protein Changes in Penetrating and Non-penetrating Models of Traumatic Brain Injury. Military Health System Research Symposium, Young Investigators' Forum.
13) PROFESSIONAL AWARDS RECEIVED DURING TENURE
N/A
14) POST-TENURE POSITION / JOB TITLE
Research Biologist
15) NAME AND ADDRESS OF POST-TENURE POSITION / JOB ORGANIZATION
Walter Reed Army Inst. For Research
16) POST-TENURE POSITION STATUS / CATEGORY Please indicate only one.  ☐ Permanent position at the NRC host agency ☐ Research/administration position in private industry in the U.S. ☐ Contract or temporary position at the NRC host Agency Abbreviate Host Laboratory/Center the U.S. ☐ Research/Administrative position with another U.S ☐ Research/administration position with a non profit
government agency  Research/Administrative position with a foreign-  Self-employed/consulting  Postdoctoral research

☐ No information provided

Other (Please specify, possible)

17) APPRAISAL OF RESEARCH ASSOCIATESHIP PROGRAM

On a scale of 1-10 (poor - excellent), please rate the following:

Research/teaching position at a U.S. college or university
Research/teaching position at a foreign college or university

**SHORT TERM VALUE** 

government agency

Development of knowledge, skills, and research productivity

#### **Comments**

#### LONG TERM VALUE

10 How the NRC Associateship award affected your career to date

**Comments** 

#### LAB SUPPORT

Quality of support from the Laboratory--equipment, funding, orientation, safety and health guidelines, etc.

**Comments** 

#### ADVISER/MENTOR SUPPORT

Quality of mentoring from the Laboratory NRC Adviser (USMA Mentor, if applicable)

**Comments** 

#### LPR SUPPORT

Quality of administrative support from the <u>L</u>aboratory (e.g., NIST, NRL, IWR, FHWA) NRC <u>P</u>rogram <u>R</u>epresentative (LPR) Comments

#### NRC SUPPORT

Quality of administrative support. Please assess respective NRC aspects (e.g., moving company, insurance, Omega, payroll, coordinator, travel, etc.)

**Comments** 

18) PLEASE PROVIDE ANY SUGGESTIONS FOR PROGRAM IMPROVEMENT.

Please do NOT scan to PDF. Send the Final Report as MSWord document via e-mail to your NRC Program Coordinator

No handwritten signature required; but you may upload a scanned signature file below: Maria Crocco: mcrocco@nas.edu
Asha Davis: adavis@nas.edu
Linda Sligh: lsligh@nas.edu
Jason Thornhill: jthornhill@nas.edu
Peggy Wilson: pwilson@nas.edu

Id# Rev. Jan 2013 Proj/Act ID#

Advisers to the Nation on Science, Engineering, and Medicine National Research Council

#### **Research Associateship Programs**

#### FINAL REPORT

1) <i>Ass</i>	ociate Last or Family Na	me	First Name		M.I.
Chen			Guojun		
3) <i>Tod</i>	'ay's Date		Dates of Tenure		
April	18, 2012		from <b>08-08-2011</b>	to August 8, 2012	
4)	Host Agency	Laboratory or Center		Division / Directorate / Departmer	ıt
	USAMC				
_	(e.g., AFRL)	(e.g., Wright Patterson AFB)	)	(e.g., High-Speed Propulsion)	
5) <i>Nan</i>	ne of Laboratory NRC Ad	viser (and USMA Mentor, if applicable)			
Da	avid. Devore				

- 6) TITLE OF RESEARCH PROPOSAL
- 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.
  - 1) virus-like particle for antisense ODN delivery
  - 2) peptide-oligonucleic acid conjugates for antisense
  - 3) Co-polymer enhanced liposome for antisense ODN delivery
  - 4)
  - 5)
- (USMA Davies Fellow: please add summary of teaching, including classes taught.)
- 8) RESEARCH IN PROGRESS Describe in no more than 100 words.

Virus-like particle project was stopped due to the lack of fresh plant infected tissue. Therefore, two extra backup plans were carried out. One is peptide-oligonucleic acid (PNA) and the other one is liposome delivery system. For PNA project, conjugation are being synthesized and characterized. The preliminary bacteria tests have indicated that PNA can improve the antisense antimicrobial capability. For copolymer enhanced liposome, the copolymers were synthesized. However, the complexing the biological testing are not carried out yet.

9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.

- a) Publications in peer-reviewed journals
- b) Books, book chapters, other publications
- c) Manuscripts in preparation, manuscripts submitted
  - Peptide-oligonucleic acid for mecA antisense in S.aureus
- 10) PATENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH Provide titles, inventors, and dates of applications.
- 11) PRESENTATIONS AT SCIENTIFIC MEETINGS OR CONFERENCES

Provide complete references: author(s), title, abstract/proceeding citation, meeting name and location.

International

Don	nestic
12) <i>SEl</i>	MINARS OR LECTURES DELIVERED AT UNIVERSITIES AND/OR INSTITUTES Include dates, names and locations of seminars.
13) <i>PR</i> (	OFESSIONAL AWARDS RECEIVED DURING TENURE
14) <i>POS</i>	ST-TENURE POSITION / JOB TITLE
En  16) POS  Peri Con Abb Res gov	mory/Atlanta  ST-TENURE POSITION STATUS / CATEGORY Please indicate only one. manent position at the NRC host agency Research/administration position in private industry in the U.S. market or temporary position at the NRC host Agency Research/administration position in private industry outside of the U.S. mearch/Administrative position with another U.S mearch/Administrative position with a foreign-  ME AND ADDRESS OF POST-TENURE POSITION / JOB ORGANIZATION  Research/administration position in private industry outside of the U.S. Research/administration position with a non profit Self-employed/consulting  Postdoctoral research
Res	rernment agency earch/teaching position at a U.S. college or university earch/teaching position at a foreign college or university  Other (Please specify, possible)  No information provided
On a s	PRAISAL OF RESEARCH ASSOCIATESHIP PROGRAM scale of 1 – 10 (poor - excellent), please rate the following:  HORT TERM VALUE  Development of knowledge, skills, and research productivity  Comments
<u>LC</u>	ONG TERM VALUE  How the NRC Associateship award affected your career to date  Comments
<u>LA</u>	AB SUPPORT  Quality of support from the Laboratoryequipment, funding, orientation, safety and health guidelines, etc.

#### Comments

#### ADVISER/MENTOR SUPPORT

Quality of mentoring from the Laboratory NRC Adviser (USMA Mentor, if applicable)

Comments

#### LPR SUPPORT

Quality of administrative support from the <u>L</u>aboratory (e.g., NIST, NRL, IWR, FHWA) NRC <u>P</u>rogram <u>R</u>epresentative (LPR) Comments

#### <u>NRC SUPPORT</u>

Quality of administrative support. Please assess respective NRC aspects (e.g., moving company, insurance, Omega, payroll, coordinator, travel, etc.)

**Comments** 

18) PLEASE PROVIDE ANY SUGGESTIONS FOR PROGRAM IMPROVEMENT.

 $signature\ file\ below: \\ \textbf{Jason\ Thornhill:}\ \underline{\textbf{jthornhill@nas.edu}}$ 

Peggy Wilson: pwilson@nas.edu
Suzanne White: swhite@nas.edu

Id# Rev. July 2011 Proj/Act ID#

Advisers to the Nation on Science, Engineering, and Medicine National Research Council

#### **Research Associateship Programs**

#### FINAL REPORT

1) Associate Last or Family Nam	ne	First Name	I I	M.I.
Johnston		Sara		C
3) Today's Date		Dates of Tenure		
April 26, 2012		from March 25,	2009 to April 30, 2012	
4) Host Agency	Laboratory or Center		Division / Directorate / Department	
USAMRIID	Fort Detrick Army Garri	iso	Virology Division	
(e.g., AFRL)	(e.g., Wright Patterson AFB)	)	(e.g., High-Speed Propulsion)	
5) Name of Laboratory NRC Adv	iser (and USMA Mentor, if applicable)			
Dr. Arthur Goff				

6) TITLE OF RESEARCH PROPOSAL

Identification and Characterization of Viral Immunomodulators that Affect the Host Specificity of Orthopoxviruses

- 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.
  - 1) MPXV Active Disease Surveillance Program identifying significant increases in prevalence in DRC
  - 2) Identification of potential novel therapeutic (IFN-beta) against MPXV
  - 3) Identification of MPXV variants actively circulating in the DRC
  - 4) In vitro and in vivo evaluation of novel therapeutics against Henipaviruses
- 5) Development and implementation of diagnostic ELISA assays against MPXV (USMA Davies Fellow: please add summary of teaching, including classes taught.)
- 8) RESEARCH IN PROGRESS Describe in no more than 100 words.

Continued efforts associated with MPXV active disease surveillance which are ongoing, in vivo characterization of IFN-beta against MPXV, continued therapeutic testing against Henipaviruses.

#### 9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.

- a) Publications in peer-reviewed journals
  - "In vitro inhibition of monkeypox virus production and spread by Interferon-beta" Sara C Johnston, Kenny L Lin, John H Connor, Gordon Ruthel, Arthur Goff, and Lisa E Hensley: Virology Journal 2012, 9:5
  - "Assessment of high-throughput screening (HTS) methods for high-consequence pathogens" Brian M Friedrich, Corinne E Scully, Jennifer M Brannan, Monica M Ogg, Sara C Johnston, Lisa E Hensley, Gene G Olinger, and Darci R Smith: Bioterrorism & Biodefense 2011, S3
  - "Development of a novel nonhuman primate model for Rift Valley Fever" Darci R Smith, Brian H Bird, Bridget Lewis, Sara C Johnston, Sarah McCarthy, Ashley Keeney, Miriam Botto, Ginger Donnelly, Joshua Shamblin, Cesar G Albarino, Stuart T Nichol, and Lisa E Hensley: Journal of Virology 2012, 86(4):2109-2120
  - "Using remote sensing to map the risk of human monkeypox virus in the Congo basin" Trevon Fuller, Henri A Thomassen, Prime M Mulembakani, Sara C Johnston, James O Lloyd-Smith, Neville K Kisalu, Timothee K Lutete, Seth Blumberg, Joseph N Fair, Nathan D Wolfe, Robert L Shongo, Pierre Formenty, Hermann Meyer, Linda L Wright, Jean-Jacques Muyembe, Wolfgang Buermann, Sassan S Saatchi, Emile Okitolonda, Lisa Hensley, Thomas B Smith, and Anne W Rimoin: EcoHealth 2010, 8(1):14-25
  - "Major increase in human monkeypox incidence 30 years after smallpox vaccination campaigns cease in the Democratic Republic of Congo" Anne W Rimoin, Prime M Mulembakani, Sara C Johnston, James O Lloyd Smith, Neville K Kisalu, Timothee L Kinkela, Seth Blumberg, Henri A Thomassen, Brian L Pike, Joseph N Fair, Nathan D Wolfe, Robert L

Shongo, Barney S Graham, Pierre Formenty, Emile Okitolonda, Lisa E Hensley, Hermann Meyer, Linda L Wright, and Jean-Jacques Muyembe: PNAS 2010, 107(37):16262-16267

- b) Books, book chapters, other publications
- c) Manuscripts in preparation, manuscripts submitted
  - "Risk Factors for Human Monkeypox in the Democratic Republic of the Congo"

Manuscript in Review (Emerging Infectious Diseases).

• "Identification of Genomic Destabilization in Monkeypox Clinical Samples"

**Manuscript in Preparation (Science)** 

- "Pathogen-host Associations and Range Shifts of Human Monkeypox in Response to Climate Change in Central Africa" Manuscript in Preparation (PLOS One).
- 10) PATENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide titles, inventors, and dates of applications.

#### 11) PRESENTATIONS AT SCIENTIFIC MEETINGS OR CONFERENCES

Provide complete references: author(s), title, abstract/proceeding citation, meeting name and location.

#### International

• "A Major Increase in the Incidence of Human Monkeypox Thirty Years After Smallpox Vaccination Campaigns Cease in the Democratic Republic of Congo"

Oral presentation at the 11th Meeting of the WHO Advisory Committee on Variola Virus, Geneva, Switzerland November 2009

#### **Domestic**

"Human Monkeypox Emergence Since the Cessation of Global Smallpox Vaccination"
 Oral presentation at the Pennsylvania Vector Control Association Training Conference,
 State College, PA

November 2011

- "Evaluation of Rash Illness in DRC Using Pan-Orthopox, Monkeypox Specific, and VZV Specific Assays"
   Poster presentation at the Chemical and Biological Defense Science and Technology Conference, Orlando, FL
   November 2010
- "Major Increase in Human Monkeypox Incidence Thirty Years After Smallpox Vaccination Campaigns Cease in the DRC"

Poster presentation at the XVIII International Poxvirus, Asfivirus, and Iridovirus Symposium, Sedona, AZ June 2010

- "Evaluation of Rash Illness in DRC Using Pan-Orthopox, Monkeypox Specific, and VZV Specific Assays"
   Poster presentation at the XVIII International Poxvirus, Asfivirus, and Iridovirus Symposium, Sedona, AZ
   June 2010
- "Vaccinia Virus E3L Blocks the Formation of Stress Granule (SG)-Related "Factory Granules" that Inhibit the Replication of a  $\Delta$ E3L Virus"

Oral presentation at the XVIII International Poxvirus, Asfivirus, and Iridovirus Symposium, Sedona, AZ June 2010

"The Actin Motor Myosin V Associates with Intracellular Enveloped Virions"
 Oral presentation at the XVIII International Poxvirus, Asfivirus, and Iridovirus Symposium, Sedona, AZ June 2010

"Human Monkeypox Genomic Divergence and Determinants of Pathogenesis"

Poster presentation at the 58th annual meeting of the American Society of Tropical Medicine

Washington, DC

November 2009

12) SEMINARS OR LECTURES DELIVERED AT UNIVERSITIES AND/OR INSTITUTES Include dates, names and locations of seminars.

"The Development of Antiviral Strategies to Combat Emerging/Re-Emerging Pathogens"
Oral presentation at Tufts University Cummings School of Veterinary Medicine
North Grafton, MA
April 2012

#### $Research\ Microbiologist\ 4\ contracted\ through\ ClinRM$

15) NAME AND ADDRESS OF POST-TENURE POSITION / JOB ORGANIZATION

USAMRIID. 1425 Porter St. Fort Detrick. MD 217	rter St. Fort Detrick, MD 2170	St.	1425 Porter	SAMRIID	US
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Permanent position at the NRC hos     Permanent position at the NRC hos     Contract or temporary position at the Abbreviate Host Laboratory/Center     Research/Administrative position of government agency     Research/Administrative position of government agency     Research/teaching position at a U.S.     Research/teaching position at a formula of the properties of	t agency ne NRC host Agency with another U.S with a foreign- S. college or university	Research/administration Research/administration the U.S.	le)
17) APPRAISAL OF RESEARCH ASSOCIATION of a scale of 1 – 10 (poor - excellent SHORT TERM VALUE  Development of knowledge, second comments	nt), please rate the followi		
LONG TERM VALUE How the NRC Associateship a Comments	award affected your career	to date	
LAB SUPPORT Quality of support from the L Comments	aboratoryequipment, fun	ding, orientation, safety and heal	th guidelines, etc.
ADVISER/MENTOR SUPPORT Quality of mentoring from the Comments		(USMA Mentor, if applicable)	
LPR SUPPORT Quality of administrative supports	port from the $\underline{\mathbf{L}}$ aboratory ( $\epsilon$	e.g., NIST, NRL, IWR, FHWA)	NRC <u>Program</u> <u>Representative</u> (LPR)
NRC SUPPORT Quality of administrative support coordinator, travel, etc.) Comments	port. Please assess respecti	ve NRC aspects (e.g., moving co	ompany, insurance, Omega, payroll,
18) PLEASE PROVIDE ANY SUGGESTIO	NS FOR PROGRAM IMPRO	VEMENT.	
but you may upload a scanned signature file below:	Asha Davis: adavis@ Linda Sligh: Isligh@r Jason Thornhill: jthornhi	nas.edu as.edu ll@nas.edu @nas.edu	our NRC Program Coordinator
Id#	Rev.	July 2011	Proj/Act ID#

Advisers to the Nation on Science, Engineering, and Medicine National Research Council

#### **Research Associateship Programs**

#### FINAL REPORT

1) Associate Last or Family Name			First Name		M.I.
Langer			Thomas		
3) <i>To</i>	oday's Date		Dates of Tenure		
novembre 15, 2012			from agosto 28, 2	2011 to novembre 23, 2012	2
4)	Host Agency	Laboratory or Center		Division / Directorate / Departme	nt
	AMRMC	USA ISR			
(e.g., AFRL) (e.g., Wright Patterson AFB			(e.g., High-Speed Propulsion)		
5) Name of Laboratory NRC Adviser (and USMA Mentor, if applicable)					
A	Andriy Batchinsky and Leopoldo Cancio				

6) TITLE OF RESEARCH PROPOSAL

PHYSIOLOGY AND PATHOPHYSIOLOGY OF SPONTANEOUS BREATHING DURING TOTAL EXTRACORPOREAL RESPIRATORY SUPPORT IN HEALTHY SHEEP AND IN SHEEP WITH THE ACUTE RESPIRATORY DISTRESS SYNDROME

- 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.
  - 1) Development of a large animal model to study spontaneous breathing during extracorporeal gas exchange
  - 2) Study of differences in respiratory pattern between healthy animals and animals with the Acute Respiratory Distress Syndrome
  - 3) Development of a new techique for the placement of bicaval dual-lumen catheters for venovenous extracorporeal gas exchange
  - 4) Effects of radiation dose reduction on lung quantitative CT scan results in healthy in the Acute Respiratory Distress Syndrome: low-dose chest CT as a valuable tool for quantification and monitoring of pulmonary disease reducing patient exposure

5

(USMA Davies Fellow: please add summary of teaching, including classes taught.)

8) RESEARCH IN PROGRESS Describe in no more than 100 words.

Mechanical ventilation (MV) is the current standard of care for the treatment of the acute respiratory distress syndrome. MV can however worsen lung injury. Extracorporeal Gas Exchange is a tempting alternative to treat ARDS. We have established a model to start investigating this treatment option in order to be able to provide, in the future, a safe and successful treatment to patients with ARDS.

9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.

a) Publications in peer-reviewed journals

Pressure-guided positioning of bicaval dual-lumen catheters for venovenous extracorpoeral gas exchange. Intensive Care Med. 2012 Nov 17. [Epub ahead of print]

b) Books, book chapters, other publications

0

c) Manuscripts in preparation, manuscripts submitted

Extracorporeal Gas Exchange in awake spontaneously breathing sheep before and after the induction of ARDS - manuscript in preparation

Vecchi V, Langer T, Bellomi M, Rampinelli C, Chung KK, Cancio LC and Batchinsky AI. Ultra Low-dose Quantitative CT: a Monitoring Tool for the Acute Respiratory Distress Syndrome? - manuscript submitted

10)	PATENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH Provide titles, inventors, and dates of applications.  N/A
11)	PRESENTATIONS AT SCIENTIFIC MEETINGS OR CONFERENCES Provide complete references: author(s), title, abstract/proceeding citation, meeting name and location.  International
	Vecchi V, Langer T, Batchinsky AI, Ivey K, Walker K, Necsoiu C, Belenkiy SM, Dimitri R, Lucas M, Leon A, Salinas J, Cancio LC. Semi-automated Quantitative CT-scan Analysis For In Vivo Lung Weight Measurement. Experimental Biology 2012, San Diego, CA.
	Domestic
	Vecchi V, Langer T, Batchinsky AI, Ivey K, Walker K, Necsoiu C, Belenkiy SM, Dimitri R, Lucas M, Leon A, Meyers B, Salinas J, Cancio LC. Semi-automated Quantitative CT-scan Analysis For In Vivo Lung Weight Measurement. Military Health System Research Symposium 2012, Ft Lauderdale, Florida.
12)	) SEMINARS OR LECTURES DELIVERED AT UNIVERSITIES AND/OR INSTITUTES Include dates, names and locations of seminars 0
13)	) PROFESSIONAL AWARDS RECEIVED DURING TENURE  0
14)	POST-TENURE POSITION / JOB TITLE
	Intensice Care Medicine Resident
15)	NAME AND ADDRESS OF POST-TENURE POSITION / JOB ORGANIZATION
	Istituto di Anestesia e Rianimazione, Università degli Studi di Milano, Milan, Italy
	POST-TENURE POSITION STATUS / CATEGORY Please indicate only one.  Permanent position at the NRC host agency
	n a scale of 1 – 10 (poor - excellent), please rate the following:
	SHORT TERM VALUE Development of knowledge, skills, and research productivity Comments Great opportunity to develop research skills  LONG TERM VALUE How the NRC Associateship award affected your career to date Comments
	<ul> <li>LAB SUPPORT</li> <li>Quality of support from the Laboratoryequipment, funding, orientation, safety and health guidelines, etc.</li> <li>Comments</li> </ul>
	ADVISER/MENTOR SUPPORT Quality of mentoring from the Laboratory NRC Adviser (USMA Mentor, if applicable) Comments
	<u>LPR SUPPORT</u>

Quality of administrative support from the <u>L</u>aboratory (e.g., NIST, NRL, IWR, FHWA) NRC <u>Program</u> <u>Representative</u> (LPR)

#### **Comments**

#### NRC SUPPORT

Quality of administrative support. Please assess respective NRC aspects (e.g., moving company, insurance, Omega, payroll, coordinator, travel, etc.)

#### **Comments**

Id#

Really great support. Thank you Jason and Peggy!

18) PLEASE PROVIDE ANY SUGGESTIONS FOR PROGRAM IMPROVEMENT.

Please do NOT scan to PDF. Send the Final Report as MSWord document via e-mail to your NRC Program Coordinator

No handwritten signature required; but you may upload a scanned signature file below:

Linda Sligh: sligh@nas.edu

Jason Thornhill: jthornhill@nas.edu

Peggy Wilson: pwilson@nas.edu

Suzanne White: swhite@nas.edu

Rev. July 2011 Proj/Act ID#

Advisers to the Nation on Science, Engineering, and Medicine National Research Council

#### **Research Associateship Programs**

## FINAL REPORT

1) Associate Last or Family Name			First Name		M.I.
Leung			Lai Yee		
3) Today's Date			Dates of Tenure		
February 1, 2013			from February 15, 2010 to February 15, 2013		
4)	Host Agency	Laboratory or Center		Division / Directorate / Departme	ent
	AMRMC	WRAIR		PNS	
(e.g., AFRL) (e.g., Wright Patterson AFB			B) (e.g., High-Speed Propulsion)		
5) Name of Laboratory NRC Adviser (and USMA Mentor, if applicable)					
Frank Tortella					

6) TITLE OF RESEARCH PROPOSAL

Characterizing a clinically/militarily relevant rat model of polytrauma associated with penetrating brain injury: neuropathological effects of transient hypotension and/or acute hypoxemia after penetrating ballistic-like brain injury (PBBI)

- 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.
  - 1) Established polytrauma models associated with hypoxemia and hemorrhagic hypotension that will be used for future neuroprotective drug studies.
  - 2) Acute physiological changes were characterized in the polytrauma models. The patterns of these changes were found to be unique under different injury combinations.
  - 3) Hemorrhagic shock increased the incidence and duration of cortical spreading depolarization within 2 hours following PBBI whereas hypoxemia only prolonged the depolarization.
  - 4) Histopathological changes were characterized (3, 7 days post-injury) in the polytrauma models. Hemorrhagic shock increased neuronal degeneration and astrocytic activation following PBBI.
  - 5) The sequence of PBBI, hemorrhagic hypotension and hypoxemia affected the mortality rate and neurological deficits. PBBI, HS followed by HX resulted in the highest mortality rate (50%) and more neurological deficits among all polytrauma groups.

(USMA Davies Fellow: please add summary of teaching, including classes taught.)

#### 8) RESEARCH IN PROGRESS Describe in no more than 100 words.

Part of the histopathological characterization is underway. Brain samples collected following the different sequences of polytrauma are being processed for immunostaining. The quantifications of histopathological changes will be performed in these immunotained brain slices. The behavioral and EEG studies in the polytrauma models will soon be started. I have submitted a core funding proposal (FY14-18) as a principal investigator to Combat Casualty Care Research Program of MRMC to continue the research efforts on polytrauma. The proposal aims at investigating the effects of polytrauma on protein biomarkers, the vital organs, etc. as well as polytrauma associated with mild TBI.

#### 9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.

a) Publications in peer-reviewed journals

Chen Z\*, Leung LY\*, Mountney A, Liao Z, Yang W, Lu XC, Dave JR, Deng-Bryant Y, Wei G, Schmid K, Shear DA, Tortella FC. A novel animal model of closed-head concussive-induced mild traumatic brain injury: development, implementation and characterization. J Neurotrauma. 2012 Jan 20;29(2):268-80. (\*Co-first authors)

Murakami Y, Wei G, Yang X, Lu XC, Leung LY, Shear DA, Tortella FC. Brain oxygen tension monitoring following penetrating ballistic-like brain injury in rats. J Neurosci Methods. 2012 Jan 15;203(1):115-21.

Mountney A\*, Leung LY\*, Pedersen R, Shear DA, Tortella FC. Longitudinal assessment of gait abnormalities following penetrating ballistic-like brain injury in rats. J Neurosci Methods. 2013 Jan 15, 212(1): 1–16. (\*Co-first authors)

Leung LY, Wei G, Shear DA, Tortella FC. The acute effect of hemorrhagic shock on cerebral blood flow, brain tissue oxygen tension and spreading depolarization following penetrating ballistic-like brain injury. Journal of Neurotrauma. (Final revision)

Lu XCM, Mountney A, Chen Z, Wei G, Leung LY, Cao Y, Khatri V, Cunningham T, Tortella FC. Similarities and differences of acute non-convulsive epileptic activities following penetrating and ischemic brain injury in rats. Journal of Neurotrauma. (In Press)

- b) Books, book chapters, other publications
- c) Manuscripts in preparation, manuscripts submitted

Leung LY, Wei G, Shear DA, Tortella FC. Temporal and Spatial Profile of Histopathological Changes Caused by Hemorrhagic Shock in a Rat Model of Penetrating Ballistic-like Brain Injury (PBBI). (In preparation)

10) PATENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH Provide titles, inventors, and dates of applications.

N/A

11) PRESENTATIONS AT SCIENTIFIC MEETINGS OR CONFERENCES

Provide complete references: author(s), title, abstract/proceeding citation, meeting name and location.

International

N/A

**Domestic** 

Leung LY, Wei G, Shear DA, Tortella FC. Acute effect of transient hypoxemia on cerebral blood flow, brain tissue oxygen tension and direct-current electroencephalography following penetrating ballistic-like brain injury. Society for Neuroscience 2012. New Orleans, Louisiana, USA. 2012.

Leung LY, Wei G, Shear DA, Tortella FC. Characterization of a military-televant polytrauma model: combined effects of penetrating ballistic-like brain injury and hemorrhagic shock or hypoxemia. Military Health System Research Symposium 2012. Fort Lauderdale, Florida. 2012.

Leung LY, Larimore Z, Holmes L, McLoughlin S, Mountney A, Schmid K, Shear DA, Tortella FC. WRAIR Projectile Concussive Impact (PCI) model: injury device and helmet advanced development. Military Health System Research Symposium 2012. Fort Lauderdale, Florida. 2012. (Oral Presentation)

Leung LY, McLoughlin S, Wei G, Shear DA, Tortella FC. Temporal and spatial profile of histopathological changes caused by hemorrhagic shock after penetrating ballistic-like brain injury (PBBI). National Neurotrauma Symposium 2012. Phoenix, Arizona, USA. July 2012.

Leung LY, Wei G, Khatri V, Shear DA, Tortella FC. Development of a military-relevant polytrauma model: combined effects of penetrating ballistic-like brain injury and hemorrhagic hypotension in rats. National Neurotrauma Symposium 2011. Hollywood, Florida, USA. July 2011.

Leung LY, Chen Z, Liao Z, Lu X, Dave J, Wei G, Yang W, Schmid K, Shear DA, Tortella FC. The WRAIR Model of Projectile Concussive Impact (PCI): I. Device development. National Neurotrauma Symposium 2011. Hollywood, Florida, USA. July 2011.

Leung LY, Wei G, Khatri V, Shear DA, Tortella FC. Effects of hypotension on spreading depolarization in the rat penetrating ballistic-like brain injury model. National Capital Region Traumatic Brain Injury Research Symposium 2011. Gaithersburg, Maryland, USA. April 2011. (Oral Presentation)

- 12) SEMINARS OR LECTURES DELIVERED AT UNIVERSITIES AND/OR INSTITUTES Include dates, names and locations of seminars. 8 November, 2011. CNS Injury Conference, hosted by Center for Brain Injury and Repair, University of Pennsylvannia.
- 13) PROFESSIONAL AWARDS RECEIVED DURING TENURE

N/A

15) NAME AND ADDRESS OF POST-TENURE POSITION/JOB ORGANIZATION

Walter Reed Army Institute of Research 503 Robert Grant Avenue, 2W12 Silver Spring, Maryland

16) POST-TENURE POSITION STATUS	/ CATEGORY Ple	ease indicate o	only one.		
Permanent position at the NRC hos	st agency		Research/administ	ration position in private	industry in the U.S.
Contract or temporary position at the		icy	Research/administ	ration position in private	industry outside of
Abbreviate Host Laboratory/Center WI			the U.S.		
Research/Administrative position	with another U.S	-	Research/administ	ration position with a non	ı profit
government agency			Self-employed/cor	ısulting	
Research/Administrative position	with a foreign-		Postdoctoral resea	rch	
government agency			Other (Please specify	, possible)	
Research/teaching position at a U.			No information pr		
Research/teaching position at a for	reign college or un	niversity	_		
17) APPRAISAL OF RESEARCH ASSOCIATION 17) APPRAISAL OF RESEARCH ASSOCIATION 17 OF 18	nt), please rate the	e following:	7		
LONG TERM VALUE  How the NRC Associateship  Comments	award affected you	ur career to d	ate		
LAB SUPPORT Quality of support from the L Comments	aboratoryequipm	nent, funding	, orientation, safety an	d health guidelines, etc.	
ADVISER/MENTOR SUPPORT  Quality of mentoring from the  Comments		Adviser (US	SMA Mentor, if applic	able)	
LPR SUPPORT  Quality of administrative supports	port from the <u>L</u> abo	oratory (e.g.,	NIST, NRL, IWR, FF	IWA) NRC <u>P</u> rogram <u>R</u> epi	resentative (LPR)
NRC SUPPORT Quality of administrative support coordinator, travel, etc.) Comments	port. Please assess	respective N	RC aspects (e.g., mov	ing company, insurance, (	Omega, payroll,
18) PLEASE PROVIDE ANY SUGGESTIC	ONS FOR PROGRAM	1 IMPROVEM	IENT.		
Please do NOT scan to PDF. Send to handwritten signature required; but you may upload a scanned signature file below:	Asha Davis: Linda Sligh: Jason Thornhill: Peggy Wilson:	as MSWord adavis@nas. lsligh@nas.e jthornhill@n pwilson@na swhite@nas.	<u>edu</u> <u>du</u> 1as.edu s.edu	ail to your NRC Program	n Coordinator
		Rev. July			
Id#		Kev. July	2011	Proj/Act ID#	

Advisers to the Nation on Science, Engineering, and Medicine National Research Council

## **Research Associateship Programs**

#### FINAL REPORT

1) Associate Last or Family Name			First Name		M.I.
McC	McCoy				E
3) <i>Toa</i>	lay's Date		Dates of Tenure		
September 4, 2012			from July 29, 2012 to October 4, 2012		
4)	Host Agency	Laboratory or Center		Division / Directorate / Departmer	ıt
WRAIR David E. Lanar		Malaria Vaccine Branch			
(e.g., AFRL) (e.g., Wright Patterson AFB			(e.g., High-Speed Propulsion)		
5) <i>Nar</i>	5) Name of Laboratory NRC Adviser (and USMA Mentor, if applicable)				
D	David. E. Lanar N/A				

6) TITLE OF RESEARCH PROPOSAL

Examination of immune responses and efficacy in mouse and Rhesus animal models of a circumsporozoite proteine (CSP)-based Self-Assembling Polypeptide Nanoparticle (SAPN) malaria vaccine.

- 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.
  - 1) Identified the mechanism of action of SAPN-induced Ab that provides sterile immunity in mice
  - 2) designed and carried out experiments to examine and characterize the processing and presentation of SAPN within the immune system
  - 3) Examined the phenotypes of SAPN-specific T-cell populations and proved that CD8+ T-cells from mice immunized with SAPN are able to, by themselves, induce sterile immunity in mice- this is the first malaria vaccine to be able to show this.
  - 4) Examined needleless approcahes to vaccine delivery, including pleuronic lecithin organogel creams
  - 5) Identified potential effects on vaccine efficacy resulting from the addition of mosquito saliva
  - 6) The youngest investigator at the WRAIR to successfully write and complete a non-human primate trial for malaria

(USMA Davies Fellow: please add summary of teaching, including classes taught.)  $\ensuremath{N\!/\!A}$ 

8) RESEARCH IN PROGRESS Describe in no more than 100 words.

I have successfully completed all of my murine and Rhesus experiments with only minor finishing touches to be wrapped up.

9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.

- a) Publications in peer-reviewed journals
  - 1) Protective antibody and CD8+ T-cell responses to the Plasmodium falciparum protein induced by a polypeptide nanoparticle vaccine. Stephen Kaba, Margaret E. McCoy, Tais Doll, Peter Burkhard, Qin Guo, Debleena Dasgupta, Yongkun Yang, Christian Mittelholzer, Roberta Spaccapelo, Andrea Crisanti, and David E. Lanar. in press PLoS ONE September 2012
- b) Books, book chapters, other publications

N/A

- c) Manuscripts in preparation, manuscripts submitted
  - 1) Mechanisms of protective responses induced by the Plasmodium falciparum CSP-based self-assembling polypeptide nanoparticle vaccine. Margaret E. McCoy, Hannah Golden, Tais Doll, Yongkun Yang, Stephen Kaba, Peter Burkhard, and David E. Lanar. In submission Vaccine 2012

- 2) Protection efficacy of a self-assembling polypeptide nanoparticle targeting the Plasmodium falciparum circumsporozoite protein in non-human primates. Margaret E. McCoy, Hannah E. Golden, Stephen Kaba, Qin Guo, Debleena Dasgupta and David E. Lanar. In preparation 2012.
   10) PATENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH Provide titles, inventors, and dates of applications.
- 11) PRESENTATIONS AT SCIENTIFIC MEETINGS OR CONFERENCES

Provide complete references: author(s), title, abstract/proceeding citation, meeting name and location.

#### International

N/A

1) A nonadjuvanted self-assembling polypeptide nanoparticle vaccine induces a CD8+ T-cell response: Implications for a robust cellular response to malaria; Margaret E. McCoy, C Brando, SA Kaba, Q Guo, D Dasgupta, C Mittelholzer, TAPF Pimentael, Y Yang, P Burkhard and D Lanar. AAI 14<sup>th</sup> Annual International Congress of Immunology. Kobe, Japan. August 22-27 2010.

#### **Domestic**

- 1) Cross-presentation of exogenous peptides delivered in self-assembling polypeptide nanaoparticles: Implications for malaria vaccine development; Margaret E. McCoy and David E. Lanar. WRAIR, Young Investigator's Meeting. Silver Spring, MD. September 2009.
- 2) A nonadjuvanted self-assembling polypeptide nanoaprtice vaccine induces a CD8+ T-cell response: Implications for a robust cellular response to malaria. Margaret E. McCoy, C Brando, S Kaba, Q Guo, D Dasgupta, C Mittelholzer, TAPF Pimentel, P burkhard and D Lanar. National Foundation for Infectious Diseases 13<sup>th</sup> Annual Conference on Vaccine Research, Bethesda, MD. April 26-28 2010.
- 3) A self-assembling polypeptide nanoparticle vaccine is able to induce Ag-specific CD8+ T-cell responses: Implications for a multi-functional vaccine for malaria. Margaret E. McCoy, S Kaba, Q Guo, D Dasgupta, C Mittelholzer, TAPF Pimentel, Y Yang, P Burkhard and C Brando. FASEB src, Biology of the Immune System. Carefree, AZ. June 20-25 2010.
- 4) Mechanisms and ancillary modifications to delivery of a nonadjuvanted self-assembling polypeptide nanoparticle vaccine for malaria. Hannah Golden, M McCoy, C Brando, S Kaba, Q Guo, D Dasgupta, C Mittelholzer, TAPF Pimentel, Y Yang, EE Carroll, P Burkhard and D Lanar. WRAIR Student Research Symposium, Silver SPring, MD. July 2010.
- 5) Mechanisms and ancillary modifications to delivery of a nonadjuvanted self-assembling polypeptide nanoparticle vaccine for malaria. Hannah Golden, M McCoy, C Brando, S Kaba, Q Guo, D Dasgupta, C Mittelholzer, TAPF Pimentel, Y Yang, EE Carroll, P Burkhard and D Lanar. WRAIR World Malaria Day, Silver SPring, MD. April 2011.
- 6) A nanoparticle vaccine targeting plasmodium falciparum circumsporozoite protein confers protective humoral and cellular immunity; Margaret E. McCoy, SA Kaba, Q Guo, D Dasgupta, TAPF Doll, Y Yang, C Mittelholzer, R Spaccapelo, A Crisanti, P Burkhard and D Lanar. 61st Annual Meeting, ASTMH, Philadelphia, PA. December 4-8 2011.
- 12) SEMINARS OR LECTURES DELIVERED AT UNIVERSITIES AND/OR INSTITUTES Include dates, names and locations of seminars.
  - 1) Lectured at Blari High School, Silver Spring, MD 2011 and 2012
  - 2) Lectured at Bishop O'Connell High School, Arlington, VA 2011
- 13) PROFESSIONAL AWARDS RECEIVED DURING TENURE
  - 1) Young Investigator Award, 2<sup>nd</sup> runner-up, ASTMH December 2011
  - 2) The Maurice R. Hilleman Early Stage Career Investigator Award, Runner-up, the National Foundation for Infectious Diseases, April 27, 2010
- 14) POST-TENURE POSITION / JOB TITLE

Law student, the University of Texas at Austin

15) NAME AND ADDRESS OF POST-TENURE POSITION / JOB ORGANIZATION

#### Austin, TX

16) POST-TENURE POSITION STATUS / CATEGORY Please indic	ate only one.
Permanent position at the NRC host agency	Research/teaching position at a foreign college or university
Contract or temporary position at the NRC host Agency	Research/administration position in private industry in the U.S.
Abbreviate Host Laboratory/Center	Research/administration position in private industry outside of
Research/Administrative position with another U.S	the U.S.
government agency	Research/administration position with a non profit
Research/Administrative position with a foreign-	Self-employed/consulting
government agency	Postdoctoral research
Research/teaching position at a U.S. college or university	Other (Please specify, possible) <u>Law School</u>

☐ No information provided

17) APPRAISAL OF RESEARCH ASSOCIATESHIP PROGRAM

On a scale of 1 - 10 (poor - excellent), please rate the following:

#### SHORT TERM VALUE

Development Comments

Development of knowledge, skills, and research productivity

This position has truly expanded my knowledge, experience and research focusing abilities.

#### **LONG TERM VALUE**

10 How the NRC Associateship award affected your career to date

**Comments** 

This fellowship has set me apart and provided me with an outstanding base from which to launch the next phase of my career.

#### LAB SUPPORT

[10] Quality of support from the Laboratory--equipment, funding, orientation, safety and health guidelines, etc.

**Comments** 

Dr. Lanar financially supported me in more than one endeavour of mine in the lab and has encouraged my idependant thoughts on experimental designs and writing.

#### ADVISER/MENTOR SUPPORT

Quality of mentoring from the Laboratory NRC Adviser (USMA Mentor, if applicable)

**Comments** 

Dr. Lanar financially supported me in more than one endeavour of mine in the lab and has encouraged my idependant thoughts on experimental designs and writing

#### LPR SUPPORT

Quality of administrative support from the <u>L</u>aboratory (e.g., NIST, NRL, IWR, FHWA) NRC <u>P</u>rogram <u>R</u>epresentative (LPR) Comments

This position just recently changed hands, but I have to say that Dr. Sara Rothman is the most amazing mentor and support for all of us. She is an icon and a role model to which any after her can only desire to imitate.

#### NRC SUPPORT

Quality of administrative support. Please assess respective NRC aspects (e.g., moving company, insurance, Omega, payroll, coordinator, travel, etc.)

#### **Comments**

I had several major issues with payroll. Omegatravel and Jason and his crew (Ms. Winstead included) have been wonderful and very helpful whenever I needed their assiatance!!

18) PLEASE PROVIDE ANY SUGGESTIONS FOR PROGRAM IMPROVEMENT.

N/A

#### Please do NOT scan to PDF. Send the Final Report as MSWord document via e-mail to your NRC Program Coordinator

No handwritten signature required;
but you may upload a scanned
signature file below:

Linda Sligh:
Jason Thornhill:
Peggy Wilson:
Peggy Wilson:
Suzanne White:

Suzanne White:

Asha Davis:

| sdavis@nas.edu |
| sligh@nas.edu |
| pwilson@nas.edu |
| pwilson@nas.edu |
| swhite@nas.edu |

Id# Rev. July 2011 Proj/Act ID#

Advisers to the Nation on Science, Engineering, and Medicine National Research Council

## **Research Associateship Programs**

#### FINAL REPORT

1) Associate Last or Family Name			First Name		M.I.
Mel	edeo		Michael		A
3) <i>Ta</i>	oday's Date		Dates of Tenure		
July 6, 2012			from March 1, 2010 to July 6, 2012		
4)	Host Agency	Laboratory or Center		Division / Directorate / Departmen	t
	<b>AMRMC</b>	ISR		Damage Control Resuscitation	n
(e.g., AFRL) (e.g., Wright Patterson AFB)		B) (e.g., High-Speed Propulsion)			
5) Name of Laboratory NRC Adviser (and USMA Mentor, if applicable)				·	
Bowman Philip					

6) TITLE OF RESEARCH PROPOSAL

A strategy for the control of acute coagulopathy of trauma

- 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.
  - 1) Aptamers can be used to completely inhibit the anti-coagulant effects of activated protein C (aPC); however, aPC does not appear to be the sole sufficient cause of the acute coagulopathy of trauma.
  - 2) Exposing in vitro cultures of endothelial cells (ECs) to laminar flow (as a model of their physiological environment) will induce a number of changes to both EC morphology and gene expression in a variety of inflammatory and morphology pathways.
  - 3) While not all of the changes in gene expression result in an altered proteome, there are a number of significant differences in protein expression between static cultured ECs and those exposed to flow.
  - 4) An analysis of the EC glycocalyx through confocal microscopy and western blotting of membrane proteins and associated glycoforms has led to an enhanced understanding of the structure and function of the endothelial glycocalyx layer.
- 5) All of these have provided advancement in the formulation of in vitro models of endothelium; in the future it should be possible to use these models as a platform for the testing of both therapeutics and diagnostics for vascular dysfunction. (USMA Davies Fellow: please add summary of teaching, including classes taught.)

  n/a
- 8) RESEARCH IN PROGRESS Describe in no more than 100 words.

The research in progress will continue over the next few months since I will be working just down the hall from my NRC position. This includes a more thorough analysis of the proteomic effects of laminar flow on the endothelial cells through a small grant with the UTSA Proteomics Core. There is a little more work to be done with the confocal analysis of ECs particularly in a time course study of the formation of the EGL. Finally, only recently has the method for isolating and concentrating the membrane proteins for the purposes of identifying enhancement or inhibition of EGL members; more work can quickly be completed on identifying the affected members through western blotting.

#### 9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.

a) Publications in peer-reviewed journals

Bowman PD, Wang X, Meledeo MA, Dubick MA, Kheirabadi BS. Toxicity of aluminum silicates used in hemostatic dressings toward human umbilical vein endothelial cells, HeLa cells, and RAW267.4 mouse macrophages. J Trauma. 2011 Sep;71(3):727-32.

b) Books, book chapters, other publications

n/a

c) Manuscripts in preparation, manuscripts submitted

M. A. Meledeo, J. A. Bynum, J.L. Sondeen, P.D. Bowman. Characterization of the Human Umbilical Vein Endothelial Cell Glycocalyx. In preparation for submission to Amer J Phys Cell.

M. A. Meledeo, J. A. Bynum, J.L. Sondeen, P.D. Bowman. Effects of Unidirectional Laminar Flow on Human Umbilical Vein and Human Dermal Microvascular Endothelial Cell Glycocalyx. In preparation.

10) PATENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH Provide titles, inventors, and dates of applications.

Bowman PD, Meledeo MA, Campbell JE. Tissue factor-coated, biocompatible, resorbable materials for control of severe bleeding. Provisional patent application, MRMC temporary ID: ISR 11-10. Submitted Jun 1, 2011.

11) PRESENTATIONS AT SCIENTIFIC MEETINGS OR CONFERENCES

Provide complete references: author(s), title, abstract/proceeding citation, meeting name and location.

International

n/a

#### **Domestic**

Meledeo MA, Bynum JA, Sondeen JL, Bowman PD. Endothelial cell responses to laminar flow: changes in gene expression, protein, and glycocalyx. FASEB J. 2012 Mar 29; 26:1129.10. Experimental Biology 2012 conference.

Bowman PD, Bynum JA, Craig T, Meledeo MA, Darlington DT. Temporal gene expression profiling of the hypothalamus following trauma. FASEB J. 2012 Mar 29; 26:lb673. Experimental Biology 2012 conference.

Meledeo MA, Campbell JE, Bowman PD. Development of an in vitro model of the acute coagulopathy of trauma. ATACCC 2011 conference.

Meledeo MA, Campbell JE, Bowman PD. Development of an in vitro model of the acute coagulopathy of trauma. FASEB J. 2011 Mar 17; 25:616.20. Experimental Biology 2011 conference.

Meledeo MA, Bynum JA, Sondeen JL, Prince MD, Bowman PD. Investigation of genes mediating the response of human endothelial cells to steady-state laminar flow. FASEB J. 2011 Mar 17; 25:662.1. Experimental Biology 2011 conference.

12) SEMINARS OR LECTURES DELIVERED AT UNIVERSITIES AND/OR INSTITUTES Include dates, names and locations of seminars.

Meledeo MA. The role of the glycocalyx and laminar flow in maintenance of human endothelial cell function in vitro. United States Army Institute of Surgical Research scientific seminar series. 2012 May 9.

13) PROFESSIONAL AWARDS RECEIVED DURING TENURE

DMRDP FY2010 award. Proposal title: Application of Aptamer Technology for Identification and Control of the Acute Coagulopathy of Trauma, P.I. Michael Adam Meledeo, US Army Institute of Surgical Research. D61\_I\_10\_J6\_202. Award value: \$118,000.

14) POST-TENURE POSITION / JOB TITLE

Staff Scientist, Contractor with Cherokee Nation Tech.

15) NAME AND ADDRESS OF POST-TENURE POSITION / JOB ORGANIZATION

USAISR Blood Research Group 3650 Chambers Pass, Bldg 3610 Ft. Sam Houston, TX 78234

10 DOGE TENNINE DOGETION CHIMNE / CHECODY DI	. •
16) POST-TENURE POSITION STATUS / CATEGORY Please indicate	te only one.
Permanent position at the NRC host agency	Research/administration position in private industry in the U.S.
Contract or temporary position at the NRC host Agency	Research/administration position in private industry outside of
Abbreviate Host Laboratory/Center ISR	the U.S.
Research/Administrative position with another U.S	Research/administration position with a non profit
government agency	Self-employed/consulting
Research/Administrative position with a foreign-	Postdoctoral research
government agency	Other (Please specify, possible)
Research/teaching position at a U.S. college or university	No information provided
Research/teaching position at a foreign college or university	

17) APPRAISAL OF RESEARCH ASSOCIATESHIP PROGRAM

On a scale of 1-10 (poor - excellent), please rate the following:

#### SHORT TERM VALUE

Development of knowledge, skills, and research productivity

Comments

I used many things I already knew how to do in this associateship, but I did have the opportunity to learn more about WHY we do the things we do; additionally, I had the opportunity to learn several new things. Productivity was not consistently stressed.

#### LONG TERM VALUE

How the NRC Associateship award affected your career to date

Although from a scientific performance metric (that being the number of publications produced), this position was not as good for my career as it could have been (a large part of that was my own fault), it did afford me the opportunity to show my skills and efforts to others in the host institute. This directly led to my new position, and hopefully to further career advancement in the future.

#### LAB SUPPORT



Quality of support from the Laboratory--equipment, funding, orientation, safety and health guidelines, etc.

Equipment and funding are unsurpased in my experience. We had everything we needed. Safety and irrelevant training were overstressed, which is not surprising considering that those programs have to be tailored to the least common denominator.

#### ADVISER/MENTOR SUPPORT



Quality of mentoring from the Laboratory NRC Adviser (USMA Mentor, if applicable)

**Comments** 

Dr. Bowman is one of the most knowledgable people I've ever known, and he was an invaluable resource for any questions I had. He taught me a great deal about the research subjects on which I was working. My only complaints are that his expectations for my work were rarely known to me.

#### LPR SUPPORT



Quality of administrative support from the Laboratory (e.g., NIST, NRL, IWR, FHWA) NRC Program Representative (LPR) Comments

Dr. Dubick was not frequently needed or seen. He did travel a lot which made it difficult to find him, although that was a rare necessity.

#### NRC SUPPORT

Quality of administrative support. Please assess respective NRC aspects (e.g., moving company, insurance, Omega, payroll, coordinator, travel, etc.)

#### **Comments**

Administrative support was excellent. I had several problems with the travel group, but I understand that there was a lot of turmoil in that department with various people covering roles while some were out on leave. Everything worked out in the end.

#### 18) PLEASE PROVIDE ANY SUGGESTIONS FOR PROGRAM IMPROVEMENT.

My primary suggestion would be to identify to both the mentors and the associates what the NRC's expectations are for roles and performance markers. I had certain expectations (my own problem) which were not met, and by the time I realized that they were not going to be met, I felt like it was too late to say anything to anyone about it.

Please do NOT scan to PDF. Send the Final Report as MSWord document via e-mail to your NRC Program Coordinator adavis@nas.edu

No handwritten signature required; but you may upload a scanned signature file below:

Linda Sligh: Jason Thornhill: jthornhill@nas.edu Peggy Wilson: **Suzanne White:** 

**Asha Davis:** 

lsligh@nas.edu pwilson@nas.edu swhite@nas.edu

Rev. July 2011 Id#

Proj/Act ID#

Advisers to the Nation on Science, Engineering, and Medicine National Research Council

#### **Research Associateship Programs**

### FINAL REPORT

1) Associate Last or Family Name			First Name		M.I.	
Melendrez			Melanie		C	
3) Today's Date			Dates of Tenur	Dates of Tenure		
May 1, 2012			from March 2	2, 2010 to May 15, 2	2012	
4)	Host Agency	Laboratory or Cente	er	Division / Directorate /	Department	
	<b>AMRMC</b>	WRAIR		AFRIMS, Bang	gkok	
(e.g., AFRL) (e.g., Wright Patterson AFB)		AFB)	B) (e.g., High-Speed Propulsion)			
5) Name of Laboratory NRC Adviser (and USMA Mentor, if applicable)			1			
Jarman Richard G.						

6) TITLE OF RESEARCH PROPOSAL

Dengue Virus Quasispecies' role in Viral Fitness and Adaptation to Changing Environments

- 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.
  - 1) Variants analysis of dengue quasispecies populations showed that variants are host or vector specific despite containing the same consensus sequence. Diversity was found to not be constrained within the vector as suggested in some publications.
  - 2) Selection analysis showed the populations to be expanding, evolving at a faster rate than 'average' for the dataset, and were under selective pressure with a predominance of nonsynonymous mutations when compared with the consensus sequence.
  - 3) Phylogenetic analysis revealed that dengue quasispecies sequences isolated in 2010 were distinct from other ciruclating consensus sequences from Thailand and full E gene offered higher resolution than partial E gene sequences.
  - 4) Amino acid (aa) analysis suggested several positions where changes would affect replication, antibody binding or VLP assembly according to the literature. Multivariate analysis predicted uncharacterized aa positions that would have impact if altered.
- 5) This work revealed the importance of full E gene surveillance for assessment of an changes, illustrated the variability and pathogenic potential of dengue quasispecies variant diversity and established a baseline in which to make future comparisons. (USMA Davies Fellow: please add summary of teaching, including classes taught.)
- 8) RESEARCH IN PROGRESS Describe in no more than 100 words.

Future work will focus on analyzing quasispecies population dynamics over the course of an in vitro transmission cycle and characterizing changes within the population during the course of fever prior to defervescence. The merits of using cloning versus next generation sequencing technologies to assist in both of these projects will also be assessed.

#### 9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.

- a) Publications in peer-reviewed journals
  - Conlan, J.V.; R.G. Jarman, K. Vongway, P. Chinnawirotpisan, M.C. Melendrez, S. Fenwick, R.C.A. Thompson and S.D. Blacksell. 2011. Hepatitis E virus is prevalent in the pig population of Lao People's Democratic Republic and evidence exists for homogenetiv with Chinese genotype 4 human isolates. Infection, Genetics and Evolution. 11: 1306-1311.
- b) Books, book chapters, other publications
- c) Manuscripts in preparation, manuscripts submitted

Melendrez, M.C.; P. Chinnawiropisan, A. Ponlawat, C. Klungthong, S.J. Thomas, R.V. Gibbons, A.L. Rothman, T.P. Endy, I-K. Yoon, T.W. Scott, J.H. Richardson and R.G. Jarman. 2012. Quasispecies variant analysis of a 2010 dengue 3 virus from Kamphaeng Phet, Thailand. In preparation.

Melendrez, M.C.; R.G. Jarman, C. Klungthon, P. Chinnawiropisan and R.V. Gibbons. 2012. Molecular evolution of pandemic influenza A virus (pH1N1) in Nepal 2009-2010. In preparation.

Takhampunya, R.; A. Kengluecha, T. Monkanna, A. Korkusol, S. Leepitakrat, B. Tippayachai, M.C. Melendrez, B.P. Evans and J.H. Richardson. 2012. Characterization of Orientia tsutsumagushi strains isolated from Leptotrombidium mites and the rodent host post-transmission. In revision.

10) PATENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH Provide titles, inventors, and dates of applications.

11) PRESENTATIONS AT SCIENTIFIC MEETINGS OR CONFERENCES

Provide complete references: author(s), title, abstract/proceeding citation, meeting name and location.

International

MC Melendrez, P Chinnawiropisan, A Ponlawat, C Klungthong, SJ Thomas, RV Gibbons, AL Rothman, TP Endy, I-K Yoon, TW Scott, JH Richardson and RG Jarman. Quasispecies variant analysis of a 2010 dengue 3 virus from Kamphaeng Phet, Thailand. Poster Presentation. American Society of Tropical Medicine and Hygiene, Philadelphia, PA, USA. December 2011.

**Domestic** 

12) SEMINARS OR LECTURES DELIVERED AT UNIVERSITIES AND/OR INSTITUTES Include dates, names and locations of seminars.

MC Melendrez. April 26, 2012. Infectious Disease Bioinformatics: Tinker, Tailor, Soldier, Spy. USAMC-AFRIMS, Bangkok, CPB Building Room 503.

- 13) PROFESSIONAL AWARDS RECEIVED DURING TENURE
- 14) POST-TENURE POSITION / JOB TITLE

**Bioinformatics Supervisor** 

15) NAME AND ADDRESS OF POST-TENURE POSITION / JOB ORGANIZATION

Walter Reed Army Institute of Research-Viral Diseases Branch. Robert Grant Ave. Silver Spring, MD

16) POST-TENURE POSITION STATUS / CATEGORY Please indicat  ☐ Permanent position at the NRC host agency ☐ Contract or temporary position at the NRC host Agency ☐ Abbreviate Host Laboratory/Center ☐ Research/Administrative position with another U.S government agency ☐ Research/Administrative position with a foreign- government agency ☐ Research/teaching position at a U.S. college or university ☐ Research/teaching position at a foreign college or university	e only one.  Research/administration position in private industry in the U.S.  Research/administration position in private industry outside of the U.S.  Research/administration position with a non profit  Self-employed/consulting  Postdoctoral research  Other (Please specify, possible)  No information provided
	☐ No information provided

17) APPRAISAL OF RESEARCH ASSOCIATESHIP PROGRAM

On a scale of 1 - 10 (poor - excellent), please rate the following:

#### **SHORT TERM VALUE**

Development of knowledge, skills, and research productivity

**Comments** 

AFRIMS, Bangkok provided many opportunities for learning about several infectious diseases important in SE Asia. I was able to observe and participate in methodologies associated with serology (viral culturing), molecular biology (sequencing and various PCR methodologies) and field collection of mosquitoes, training provided by the Entomology Dept. at AFRIMS. The organization encouraged collaboration and laboratory space was readily available my project along with any supplies I required.

**LONG TERM VALUE** 

How the NRC Associateship award affected your career to date Comments

Having come from an environmental microbiology background with minimal training in infectious disease, tenure with NRC has encouraged me to continue in infectious disease ecology and bioinformatic analysis. It has opened opportunities for me to continue in these fields and continue working and developing under my mentors.

#### LAB SUPPORT

10 Quality of support from the Laboratory--equipment, funding, orientation, safety and health guidelines, etc.

**Comments** 

All protocols were clearly communicated and training was provided in English whenever required. Department personnel assisted me in locating and obtaining any supplies or equipment I needed for my work and clearing lab space.

#### ADVISER/MENTOR SUPPORT

Quality of mentoring from the Laboratory NRC Adviser (USMA Mentor, if applicable)

**Comments** 

Correspondence was always prompt and helpful.

#### LPR SUPPORT

Quality of administrative support from the <u>L</u>aboratory (e.g., NIST, NRL, IWR, FHWA) NRC <u>P</u>rogram <u>R</u>epresentative (LPR) Comments

Correspondence was always prompt and helpful.

#### NRC SUPPORT

Quality of administrative support. Please assess respective NRC aspects (e.g., moving company, insurance, Omega, payroll, coordinator, travel, etc.)

**Comments** 

Omega travel was very helpful with all travel arrangements. See comments below about issues pertaining to being an internationally placed postdoc.

#### 18) PLEASE PROVIDE ANY SUGGESTIONS FOR PROGRAM IMPROVEMENT.

There is not much support or guidance for internationally placed NRC postdocs except the coverage of my plane ticket. Aetna insurance also does not provide very much international support. Upon arriving to Bangkok there was no point of contact to assist in making our transition smooth. I was fortunate to have the support of the Virology secretary who served that role when I asked her to. There also was not much guidance for moving abroad specifically, I am now serving as a resource for some new NRC postdocs applying to the program as to how logistically to make moving to Bangkok Thailand feasible with respect to visas, housing, APO address, medical care, communications with NRC etc. If NRC is unable to provide this kind of abroad logistical support and the supporting agency where the fellow will work does not have this understanding to provide a point of contact for the fellow when they arrive, then the transition is challenging. Please feel free to post me as a resource to postdoctoral fellows with NRC that will be relocating abroad. I would be willing to give them information on the logistics of relocating to Thailand and advice on being an 'abroad' postdoc in general.

Please do NOT scan to PDF. Send the Final Report as MSWord document via e-mail to your NRC Program Coordinator

No handwritten signature required; but you may upload a scanned signature file below: Asha Davis: adavis@nas.edu
Linda Sligh: Isligh@nas.edu
Jason Thornhill: jthornhill@nas.edu
Peggy Wilson: pwilson@nas.edu
Suzanne White: swhite@nas.edu

Id# Rev. July 2011 Proj/Act ID#

Advisers to the Nation on Science, Engineering, and Medicine National Research Council

### **Research Associateship Programs**

### FINAL REPORT

1) Associate Last or Family Name			First Name	M.I.		
Pichugin			Alexander V			
3) Today's Date			Dates of Tenure			
July 6, 2012			from January 12, 2009 to July 11, 2012			
4)	Host Agency	Laboratory or Center		Division / Directorate / Departme	nt	
	AMRMC	WRAIR		Malaria Vaccine Branch		
(e.g., AFRL) (e.g., Wright Patterson AFB)			3) (e.g., High-Speed Propulsion)			
5) Name of Laboratory NRC Adviser (and USMA Mentor, if applicable)						
Dr. Urszula Krzych						

6) TITLE OF RESEARCH PROPOSAL

Prioritization and selection of pre-erythrocytic liver stage Plasmodia antigens as vaccine candidates.

- 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.
  - 1) 10 novel liver stage Pb antigens reduce LS and BS parasite burden in C57Bl/6 mice.
  - 2) 3 novel liver stage Pb antigens sustain protection during 6 months after the last immunization.
  - 3) 3 novel liver stage Pb antigens enhance protection induced by PbCSP.
  - 4) Established caged MHC-tetramer technology to use for discovery of T cell epitopes in malaria antigens.
  - 5) Identified 3 immunodominant CD8 T cell epitopes from Pb PEVA.

(USMA Davies Fellow: please add summary of teaching, including classes taught.)

8) RESEARCH IN PROGRESS Describe in no more than 100 words.

We showed that ten of 23 initial Pb DNA constructs induce the protection in C57Bl/6 mice after IM and Gene Gun delivery and subsequent challenge with P. berghei. Six orthologues of ten protective in P. berghei model antigens were also protective in P. yoelii-BALB/c model.

We have established novel state-of-art high throughput method of caged MHC-tetramers to evaluate frequency of peptide-specific CD8 T cells in mice protected by GAP and RAS. We have screened ~400 of Kb- and Db-restricted CD8 peptides from 28 initial Pb antigens and determined three immunodominant epitopes on two protective vaccine candidates.

9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.

- a) Publications in peer-reviewed journals
  - Anjali Yadava, Saule Nurmukhambetova, Alexander V Pichugin, Joanne M Lumsden
  - "Cross-Species Immunity Following Immunization with a Circumsporozoite Protein-Based Vaccine for Malaria" J Infect Dis. 2012, 205(9):1456-63.
- b) Books, book chapters, other publications
- c) Manuscripts in preparation, manuscripts submitted
  - "Early Liver Stage Transcriptome of Plasmodium falciparum Reveals Novel Malaria Vaccine Candidates", in preparation
- 10) PATENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH Provide titles, inventors, and dates of applications.

11) PRESENTATIONS AT SCIENTIFIC MEETINGS OR CONFERENCES Provide complete references: author(s), title, abstract/proceeding citation, meeting name and location.	
International  Alexander V. Pichugin, Lindsey Ehrler, Sharvan Sehrawat, Cate Speake, Patrick Duffy, Hidde Ploegh, Urszula Krzych  ''Application of novel caged MHC-tetramer technology for the discovery of immunodominant CD8 T cell epitopes on	
Plasmodium liver stage antigens'' 99th Annual Meeting of the American Association of Immunologists "Immunology 2012". Boston, MA, USA, May 4-8, 201 abstract 1336683, poster presentation.	2,
Alexander V. Pichugin, Cate Speake, Lindsey Ehrler, Lindsay Holladay, Valentino Garcia, Bob Morrison, Patricia DeLaVega, Isaac Chalom, D. Gray Heppner, Patrick Duffy and Urszula Krzych "Novel Plasmodium berghei pre-erythrocytic liver stage antigens as potential vaccine candidates" Keystone Symposia: Malaria: New Approaches to Understanding Host-Parasite Interactions. Cooper Mountain, Colorado, USA, April 11-16, 2010, abstract 231, poster presentation.	,
Domestic	
Alexander V. Pichugin, Stasya zarling, Lindsey Ehrler, Sharvan Sehrawat, Cate Speake, Patrick Duffy, Hidde Ploegh, Urszula Krzych "Identification of Plasmodiun berghei novel liver stage CD8 T cell epitopes by caged MHC class I tetramer technology" 2012 World Malaria Day, Johns Hopkins Malaria Research Institute. Baltimore, MD, USA, April 25, 2012, abstract 26, por presentation.	ster
12) SEMINARS OR LECTURES DELIVERED AT UNIVERSITIES AND/OR INSTITUTES Include dates, names and locations of seminary	ars.
13) PROFESSIONAL AWARDS RECEIVED DURING TENURE	
14) POST-TENURE POSITION / JOB TITLE	
Senior Immunologist	
15) NAME AND ADDRESS OF POST-TENURE POSITION / JOB ORGANIZATION	
Clinical Research Management 411 Aviation Way Ste. 220 Frederick, MD 21701	
16) POST-TENURE POSITION STATUS / CATEGORY Please indicate only one.  □ Permanent position at the NRC host agency □ Research/administration position in private industry in the U □ Research/administration position in private industry outside the U.S.  □ Research/Administrative position with another U.S□ Research/administration position in private industry outside the U.S. □ Research/Administrative position with another U.S□ Research/administration position with a non profit government agency □ Self-employed/consulting □ Postdoctoral research □ Other (Please specify, possible) □ Other (Please specify, possible) □ No information provided □ No information pro	
17) APPRAISAL OF RESEARCH ASSOCIATESHIP PROGRAM On a scale of 1 – 10 (poor - excellent), please rate the following:	
SHORT TERM VALUE  Development of knowledge, skills, and research productivity  Comments	
LONG TERM VALUE  How the NRC Associateship award affected your career to date  Comments	

LAB SUPPORT

Quality of support from the Laboratory--equipment, funding, orientation, safety and health guidelines, etc.

**Comments** 

### ADVISER/MENTOR SUPPORT

10 Quality of mentoring from the Laboratory NRC Adviser (USMA Mentor, if applicable)

**Comments** 

### LPR SUPPORT

Quality of administrative support from the <u>L</u>aboratory (e.g., NIST, NRL, IWR, FHWA) NRC <u>Program Representative (LPR)</u>
Comments

### NRC SUPPORT

Quality of administrative support. Please assess respective NRC aspects (e.g., moving company, insurance, Omega, payroll, coordinator, travel, etc.)

**Comments** 

18) PLEASE PROVIDE ANY SUGGESTIONS FOR PROGRAM IMPROVEMENT.

Please do NOT scan to PDF. Send the Final Report as MSWord document via e-mail to your NRC Program Coordinator

No handwritten signature required; but you may upload a scanned signature file below:

Asha Davis: adavis@nas.edu
Isligh@nas.edu
Jason Thornhill: jthornhill@nas.edu
Peggy Wilson: pwilson@nas.edu

Alexander Pichugin Suzanne White: <a href="mailto:swhite@nas.edu">swhite@nas.edu</a>

Id# Rev. July 2011 Proj/Act ID#

Advisers to the Nation on Science, Engineering, and Medicine National Research Council

### **Research Associateship Programs**

### FINAL REPORT

1) Associate Last or Family Nan	ne	First Name	M.I.	
Rajendran		Gnana	R	
3) Today's Date		Dates of Tenure		
September 4, 2012		from September 7, 2010 to August 31, 2012		
4) Host Agency	Laboratory or Center		Division / Directorate / Departmen	nt .
AMRMC	WRAIR		ET	
(e.g., AFRL)	(e.g., Wright Patterson AFB)	(e.g., High-Speed Propulsion)		
5) Name of Laboratory NRC Adv	iser (and USMA Mentor, if applicable)			
WRAIR DI	r. Michael Kozar			
6) TITLE OF RESEARCH PR	OPOSAL		·	
TO 1 1 11 1 0				

- Design and synthesis of decaquinate derivatives toward new antimalarial agents
- 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.
  - 1) Several compounds were synthesized by the SAR of DQ and submitted for in vitro testing against blood stage malaria, specifically P.falciparum D6, W2, C235 and C2B strains, and assessed for metabolic stability in the mouse and human microsomes
  - 2) In many cases the compounds solubility was improved but the compounds either lost potency against the C2B resistant strain of malaria or microsomal stability
  - 3) A few interesting trends were discovered. An unprecedented ester replacement, to the ethyl or morpholine amide was discovered that maintained potency against D6,W2,C235 but unfortunately lost activity against the key C2B atovaquone resistant strain
  - 4) Additionally, many compounds were synthesized that maintained in vitro potency with significantly lower clogP (main focus of the research effort)
  - 5) Efforts are currently toward acquiring a complete set of data to select profile compounds to be scaled up and tested in in vivo models

(USMA Davies Fellow: please add summary of teaching, including classes taught.)

8) RESEARCH IN PROGRESS Describe in no more than 100 words.

Please see the attached document

- 9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH
  - Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.
  - a) Publications in peer-reviewed journals
  - b) Books, book chapters, other publications
  - c) Manuscripts in preparation, manuscripts submitted
- 10) PATENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH Provide titles, inventors, and dates of applications.
- 11) PRESENTATIONS AT SCIENTIFIC MEETINGS OR CONFERENCES

International	
Domestic	
12) SEMINARS OR LECTURES DELIVERED AT UNIVERSITIES.	AND/OR INSTITUTES Include dates, names and locations of seminar
13) PROFESSIONAL AWARDS RECEIVED DURING TENURE	
14) POST-TENURE POSITION / JOB TITLE  Scientist	
15) NAME AND ADDRESS OF POST-TENURE POSITION / JOB ORGAN	TIZATION
WRAIR, 503 Robert Grant Ave, Silver Spring, MD	
16) POST-TENURE POSITION STATUS / CATEGORY Please indicated Permanent position at the NRC host agency  Contract or temporary position at the NRC host Agency Abbreviate Host Laboratory/Center WRAIR  Research/Administrative position with another U.Sgovernment agency  Research/Administrative position with a foreign-government agency  Research/teaching position at a U.S. college or university  Research/teaching position at a foreign college or university	e only one.  Research/administration position in private industry in the U.S. Research/administration position in private industry outside of the U.S. Research/administration position with a non profit Self-employed/consulting Postdoctoral research Other (Please specify, possible) No information provided
17) APPRAISAL OF RESEARCH ASSOCIATESHIP PROGRAM  On a scale of 1 – 10 (poor - excellent), please rate the following SHORT TERM VALUE  10 Development of knowledge, skills, and research productive Comments	
LONG TERM VALUE  How the NRC Associateship award affected your career to Comments	date
LAB SUPPORT  Quality of support from the Laboratoryequipment, funding Comments	ng, orientation, safety and health guidelines, etc.
ADVISER/MENTOR SUPPORT  Quality of mentoring from the Laboratory NRC Adviser (Comments	USMA Mentor, if applicable)
LPR SUPPORT  Quality of administrative support from the <u>L</u> aboratory (e.g Comments	a., NIST, NRL, IWR, FHWA) NRC <b>P</b> rogram <b>R</b> epresentative (LPR)
NRC SUPPORT Quality of administrative support. Please assess respective coordinator, travel, etc.) Comments	NRC aspects (e.g., moving company, insurance, Omega, payroll,

Please do NOT scan to PDF. Send the Final Report as MSWord document via e-mail to your NRC Program Coordinator Asha Davis: adavis@nas.edu

No handwritten signature required; but you may upload a scanned

Linda Sligh: lsligh@nas.edu signature file below: Peggy Wilson:

Jason Thornhill: <a href="mailto:jthornhill@nas.edu">jthornhill@nas.edu</a> pwilson@nas.edu **Suzanne White:** swhite@nas.edu

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Advisers to the Nation on Science, Engineering, and Medicine National Research Council

### **Research Associateship Programs**

### FINAL REPORT

1) Associate Last or Family Name			First Name		M.I.
Torres			Luciana		
3) Today's Date			Dates of Tenure		
October 18, 2012			from May 18, 2011 to July 30, 2012		
4)	Host Agency	Laboratory or Center		Division / Directorate / Departme	nt
	USAMMC	AISR		Damage Control and Resuscita	tion
	(e.g., AFRL)	(e.g., Wright Patterson AFB)	)	(e.g., High-Speed Propulsion)	
5) Name of Laboratory NRC Adviser (and USMA Mentor, if applicable)					
Dr. Michael Dubick					

- 6) TITLE OF RESEARCH PROPOSAL
- 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.
  - 1) Intravital Microscopy was successfully employed for investigating EG shedding in hemorrhagic shock/resuscitation for the very first time;
  - 2) Intravital microscopy integrated with systemic hemodynamics evaluations may be essential and more accurate tools to identify changes and study mechanisms of EG shedding and systemic responses to hemorrhage and resuscitation therapy;
  - 3) Compared to baseline and to the sham group, there was a 50% reduction in endothelial glycocalyx (EG) thickness after hemorrhage and 60% increase in the levels of plasma Syndecan-1;
  - 4) Although resuscitation with LR and Hextend could stabilize hypotensive rats hemodynamicaly, these fluids were unable to restore EG thickness or coagulopathy (weak clots and prolonged coagulation time);
  - 5) Rats who received fresh frozen plasma (FFP) restored venular EG thickness to baseline level in addition to improve the systemic hemodynamics and coagulation response (restored homeostasis).

(USMA Davies Fellow: please add summary of teaching, including classes taught.)

8) RESEARCH IN PROGRESS Describe in no more than 100 words.

We added a few more resuscitation groups (i.e. hypertonic saline, normal saline) to our protocol. So the next step is to test other resuscitation fluids and their effects on the endothelial glycocalyx (EG) degradation, coagulation and vacular permeability. Also we will continue to investigate the link between the EG and coagulopathy, in our rat hemorrhage model. Other measurements, such as vascular permeability (through fluorescence leakage) will also be started soon. A new Confocal microscope is being setup in our lab and we will be performing EG mesurements of specific componements (proteoglycans and glycosaminoglycans) in vivo and in situ in the cremaster muscle by next year.

9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.

- a) Publications in peer-reviewed journals
- b) Books, book chapters, other publications

  None
- c) Manuscripts in preparation, manuscripts submitted

Torres Filho, I.P., Torres, L.N., Sondeen, J.L., Polykratis, I.A. and Michael A. Dubick - In Vivo Evaluation of Venular Glycocalyx during Hemorrhagic Shock in Rats using Intravital Microscopy. Submitted to Microvascular Research, 2012 (under review).

Torres, L.N., Torres Filho, I.P., Sondeen, J.L., Lisa, J. and Michael A. Dubick - In Vivo Evaluation of Resuscitation Fluids on Venular Glycocalyx using Intravital Microscopy. In Preparation.
10) PATENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH Provide titles, inventors, and dates of applications.  None
11) PRESENTATIONS AT SCIENTIFIC MEETINGS OR CONFERENCES Provide complete references: author(s), title, abstract/proceeding citation, meeting name and location. International None
Domestic
Torres, L.N., Sondeen, J.L., Polykratis, I.A., Dubick, M.A. and Torres Filho I.P. – In vivo evaluation of the endothelial glycocalyst during hemorrhagic shock using intravital microscopy. SHOCK: 37, suppl 1, p.102: P223, 2012. 35th Annual Conference on Shock, Miami, Florida, 2012.
Torres, L.N., Sondeen, J.L., Ji, L., Dubick, M.A. and Torres Filho I.P. – In vivo comparison of resuscitation fluids on the preservation of the endothelial glycocalyx after hemorrhagic shock in rats. 2012 Military Health System Research Symposium (former ATACCC), Fort Lauderdale, Florida, 2012 (posters were not published in any journal supplement).
12) SEMINARS OR LECTURES DELIVERED AT UNIVERSITIES AND/OR INSTITUTES Include dates, names and locations of seminars None
12) DDGEESSIONAL AWADDS DECEMED DUDING TENUDE
13) PROFESSIONAL AWARDS RECEIVED DURING TENURE  None
14) POST-TENURE POSITION / JOB TITLE  Research Investigator
15) NAME AND ADDRESS OF POST-TENURE POSITION / JOB ORGANIZATION  US Army Institute of Surgical Research (USAISR)
16) POST-TENURE POSITION STATUS / CATEGORY Please indicate only one.  □ Permanent position at the NRC host agency □ Research/administration position in private industry in the U.S. □ Contract or temporary position at the NRC host Agency □ Research/administration position in private industry outside of the U.S. □ Research/Administrative position with another U.S□ Research/administration position with a non profit government agency □ Self-employed/consulting □ Postdoctoral research □ Other (Please specify, possible) □ No information provided □ No informatio
17) APPRAISAL OF RESEARCH ASSOCIATESHIP PROGRAM  On a scale of 1 – 10 (poor - excellent), please rate the following:  SHORT TERM VALUE  Development of knowledge, skills, and research productivity  Comments
LONG TERM VALUE  How the NRC Associateship award affected your career to date  Comments

Quality of support from the Laboratory--equipment, funding, orientation, safety and health guidelines, etc.

**Comments** 

### ADVISER/MENTOR SUPPORT

**Comments** 

Quality of mentoring from the Laboratory NRC Adviser (USMA Mentor, if applicable)

#### LPR SUPPORT

Quality of administrative support from the <u>Laboratory</u> (e.g., NIST, NRL, IWR, FHWA) NRC <u>Program</u> <u>Representative</u> (LPR) **Comments** 

#### NRC SUPPORT

Quality of administrative support. Please assess respective NRC aspects (e.g., moving company, insurance, Omega, payroll, coordinator, travel, etc.)

#### **Comments**

I could be luckier to have Jason as my coordinator: he was absolutely great, helpful and very supportive. The moving company showed efficiency, accountability, and professionalism! I am 100% satisfied. The driver was very experienced and made my move as stress-free as possible. The only complain I have is regarding the travel reimbursements: it took a long time (more than a month or 2) to receive my reimbursements, although the staff handling the travel were always very nice and knowledgeable.

18) PLEASE PROVIDE ANY SUGGESTIONS FOR PROGRAM IMPROVEMENT.

Please do NOT scan to PDF. Send the Final Report as MSWord document via e-mail to your NRC Program Coordinator

**Asha Davis:** adavis@nas.edu No handwritten signature required; lsligh@nas.edu but you may upload a scanned Linda Sligh: signature file below: Jason Thornhill: jthornhill@nas.edu pwilson@nas.edu Peggy Wilson:

**Suzanne White:** swhite@nas.edu

Rev. July 2011 Id# Proj/Act ID#

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### **Research Associateship Programs**

### FINAL REPORT

1) Associate Last or Family Name		First Name		M.I.	
Vecchi		Vittoria			
3) <i>Tod</i>	lay's Date		Dates of Tenure		
			from <b>Novemb</b>	to October 22, 2012	
4)	Host Agency	Laboratory or Center		Division / Directorate / Departme	ent
	AMRMC	USAISR			
	(e.g., AFRL)	(e.g., Wright Patterson AFB)	<u> </u>	(e.g., High-Speed Propulsion)	
5) <i>Nar</i>	ne of Laboratory NRC Advi.	ser (and USMA Mentor, if applicable)			
$\mathbf{D}_{1}$	r. Andriy Batchinsky				

6) TITLE OF RESEARCH PROPOSAL

Quantitative lung CT-scan as monitoring tool for the Acute Respiratory Distress Syndrome

- 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.
  - 1) Effects of radiation dose reduction on lung quantitative CT scan results in healthy in the Acute Respiratory Distress Syndrome: low-dose chest CT as a valuable tool for quantification and monitoring of pulmonary disease reducing patient exposure
  - 2) Use of quantitative CT for in vivo lung weight measurement: evaluate and monitor the time course of lung edema in ARDS measuring lung weight by qCT
  - 3) Pressure-guided positioning of bicaval dual-lumen catheters for veno-venous extracorporeal gas exchange
  - 4) Low-flow extracorporeal gal exchange for the treatment of ARDS caused by smoke inhalation and cutaneous burn in pigs
  - 5) Extracorporeal Gas Exchange in awake spontaneously breathing sheep before and after the induction of ARDS

(USMA Davies Fellow: please add summary of teaching, including classes taught.)  $\ensuremath{N\!/\!A}$ 

8) RESEARCH IN PROGRESS Describe in no more than 100 words.

The use of low-dose chest CT, if producing satisfactory image quality for the purpose of the quantitative analysis, could favor the use of qCT as a monitoring tool of severity and time-course of pulmonary disease reducing patient radiation exposure. The aim of the this study is therefore to investigate if and how a reduction in tube current-time product (mAs) during CT image acquisition affects quantitative results in healthy lungs and ARDS.

9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.

- a) Publications in peer-reviewed journals
- b) Books, book chapters, other publications
- c) Manuscripts in preparation, manuscripts submitted
  - Vecchi V, Langer T, Bellomi M, Rampinelli C, Chung KK, Cancio LC and Batchinsky AI. Ultra Low-dose Quantitative CT: a Monitoring Tool for the Acute Respiratory Distress Syndrome? manuscript submitted to the American Journal of Respiratory and Critical Care Medicine
  - Langer T, Vecchi V, Belenkiy SM, Cancio LC, Gattinoni L, Batchinsky AI. Pressure-guided positioning of bicaval dual-lumen catheters for veno-venous extracorporeal gas exchange. Manuscript submitted to Intensive Care Medicine
  - Extracorporeal Gas Exchange in awake spontaneously breathing sheep before and after the induction of ARDS manuscript in preparation

	TENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH wide titles, inventors, and dates of applications.
Prov	ESENTATIONS AT SCIENTIFIC MEETINGS OR CONFERENCES vide complete references: author(s), title, abstract/proceeding citation, meeting name and location. rnational
Can	chi V, Langer T, Batchinsky AI, Ivey K, Walker K, Necsoiu C, Belenkiy SM, Dimitri R, Lucas M, Leon A, Salinas J, cio LC. Semi-automated Quantitative CT-scan Analysis For In Vivo Lung Weight Measurement. Experimental Biology 2, San Diego, CA.
Dom	nestic
Saliı	chi V, Langer T, Batchinsky AI, Ivey K, Walker K, Necsoiu C, Belenkiy SM, Dimitri R, Lucas M, Leon A, Meyers B, nas J, Cancio LC. Semi-automated Quantitative CT-scan Analysis For In Vivo Lung Weight Measurement. Military lth System Research Symposium 2012, Ft Lauderdale, Florida.
12) <i>SEM</i> <b>N/A</b>	MINARS OR LECTURES DELIVERED AT UNIVERSITIES AND/OR INSTITUTES Include dates, names and locations of seminars
13) <i>PRC</i> <b>N/A</b>	OFESSIONAL AWARDS RECEIVED DURING TENURE
14) <i>POST</i>	T-TENURE POSITION / JOB TITLE
,	diology Resident
15) <i>NAM</i>	IE AND ADDRESS OF POST-TENURE POSITION / JOB ORGANIZATION
Uni	iversità degli Studi di Milano, Milan, Italy
Perm Cont Abbi Rese gove Rese gove Rese	T-TENURE POSITION STATUS / CATEGORY Please indicate only one.  nanent position at the NRC host agency
,	RAISAL OF RESEARCH ASSOCIATESHIP PROGRAM cale of 1 – 10 (poor - excellent), please rate the following:
SH0 10	ORT TERM VALUE  Development of knowledge, skills, and research productivity  Comments
<u>LO</u>	NG TERM VALUE  How the NRC Associateship award affected your career to date  Comments
LA)	B SUPPORT  Quality of support from the Laboratoryequipment, funding, orientation, safety and health guidelines, etc.  Comments
AD 10	OVISER/MENTOR SUPPORT  Quality of mentoring from the Laboratory NRC Adviser (USMA Mentor, if applicable)  Comments

### LPR SUPPORT

Quality of administrative support from the <u>Laboratory</u> (e.g., NIST, NRL, IWR, FHWA) NRC <u>Program</u> <u>Representative</u> (LPR) Comments

### NRC SUPPORT

Id#

Quality of administrative support. Please assess respective NRC aspects (e.g., moving company, insurance, Omega, payroll, coordinator, travel, etc.)

#### **Comments**

18) PLEASE PROVIDE ANY SUGGESTIONS FOR PROGRAM IMPROVEMENT.

Please do NOT scan to PDF. Send the Final Report as MSWord document via e-mail to your NRC Program Coordinator

Suzanne White: <a href="mailto:swhite@nas.edu">swhite@nas.edu</a>
Rev. July 2011
Proj/Act ID#

Advisers to the Nation on Science, Engineering, and Medicine National Research Council

## **Research Associateship Programs**

### FINAL REPORT

1) Associate Last or Family Name			First Name			M.I.
Wong			Benjamin J			J
3) Today's Date		Dates of Tenure				
			from <b>February</b>	<b>7, 2011</b>	September 4, 2012	
4)	Host Agency	Laboratory or Center		Division / Di	rectorate / Departmen	t
	MRMC	APG			MRICD	
	(e.g., AFRL)	(e.g., Wright Patterson AFB)	)	(e.g., Hig	h-Speed Propulsion)	
5) Name of Laboratory NRC Adviser (and USMA Mentor, if applicable)				·		
Dr Alfred Sciuto						

6) TITLE OF RESEARCH PROPOSAL

Treating Chemical Agent Exposure Through Multiple Drug Delivery Routes

- 7) SUMMARY OF RESEARCH DURING TENURE Itemize significant findings in concise form, utilizing key concepts/words.
  - 1) Developed and characterized novel system for inhalational exposure of conscious animals to chemical agents
  - 2) Utilized above system to examine the toxicokinetics (TK) of nerve agents and their analogs in rats
  - 3) Investigated utility of bronchodilators in a treatment regimen for inhalational chemical agent exposure
  - 4)
  - 5)

(USMA Davies Fellow: please add summary of teaching, including classes taught.)

8) RESEARCH IN PROGRESS Describe in no more than 100 words.

Current research focuses on the inhalation of various chemical agents and the associated effects on different organ systems in order to characterize the role of exposure route on progression and presence of symptoms. The body of literature that describes the inhalation of chemical agents is incomplete, despite being the most likely route by which agent will enter the body. The route of exposure is central to the progression and presence of symptoms, which in turn form the basis for the development of treatments and treatment regimens. Recent development of a novel inhalation model has progressed research toward development and characterization of improved methods of drug delivery in chemical agent exposure scenarios.

9) PUBLICATIONS AND PAPERS RESULTING FROM NRC ASSOCIATESHIP RESEARCH

Provide complete citations: author(s), title, full name of journal, volume number, page number(s), and year of publication.

a) Publications in peer-reviewed journals

N/A

b) Books, book chapters, other publications

N/A

- c) Manuscripts in preparation, manuscripts submitted
  - 1. Development of Model for Nerve Agent Inhalation in Conscious Rats
  - 2. Respiration Toxicity in Non-Anesthetized Rats Following Inhalation Exposure to Soman Vapor
  - 2 others, as yet untitled, in preparation
- 10) PATENT OR COPYRIGHT APPLICATIONS RESULTING FROM NRC ASSOCIATESHIP RESEARCH Provide titles, inventors, and dates of applications.

Provide complete references: author(s), title, abstract/proceeding citation, meeting name and location.
International
N/A
Domestic
1. B. Wong, A. Sciuto, G. Murphy. Development of a Head-Out Vapor Inhalation Model for the Evaluation of Toxicity Following Exposure to Chemical Agents in Non-Anesthetized Rats. US Army Medical Defense Bioscience Review, Hunt Valley, MD.
2. B. Wong, A. Sciuto, G. Murphy A Head-Out Vapor Inhalation Model for the Evaluation of Soman Toxicity in Non-Anesthetized Rats Shoresh Chemical and Biological Defense Conference, Fort Detrick, MD.
12) SEMINARS OR LECTURES DELIVERED AT UNIVERSITIES AND/OR INSTITUTES Include dates, names and locations of seminars.
N/A
13) PROFESSIONAL AWARDS RECEIVED DURING TENURE
N/A
14) POST-TENURE POSITION / JOB TITLE
Post-Doctoral Research Associate at MRICD
15) NAME AND ADDRESS OF POST-TENURE POSITION / JOB ORGANIZATION
USAMRICD 3100 Rickett's Point Road APG, MD 21010
16) POST-TENURE POSITION STATUS / CATEGORY Please indicate only one.  Permanent position at the NRC host agency Research/administration position in private industry in the U.S.  Contract or temporary position at the NRC host Agency Research/Administrative position with another U.S.  Research/Administrative position with another U.S  government agency Research/Administrative position with a foreign- government agency Self-employed/consulting Postdoctoral research government agency Other (Please specify, possible) Other (Please specify, possible) Research/teaching position at a foreign college or university No information provided
17) APPRAISAL OF RESEARCH ASSOCIATESHIP PROGRAM On a scale of 1 – 10 (poor - excellent), please rate the following:
SHORT TERM VALUE  Development of knowledge, skills, and research productivity  Comments  Work at MRICD was intended to continue the research from my graduate work; however, the research took on its own directions, resulting in development of knowledge and skills in new areas which complemented my existing knowledge base well. Despite many technical hurdles, challenged proved to help maximize productivity. Overall, work here has provided a different set of challenges, which have resulted in net positive gains in all categories.  LONG TERM VALUE

### LAB SUPPORT

**Comments** 

Quality of support from the Laboratory--equipment, funding, orientation, safety and health guidelines, etc.

Cannot currently ascertain effects, as a permanent position at my institute was not available, rendering it difficult to assess the competitive benefits of an NRC fellowship. However, the NRC program has helped me make connections which are

How the NRC Associateship award affected your career to date

likely to have a beneficial effect on my career in the future.

#### **Comments**

Funding has struggled for the institute recently, and orientation was somewhat confusing. Guidelines for safety and health are excellently implemented in the laboratory setting but less observed in the office setting. Host laboratory/advisor group has done the best possible given the overall situation.

### ADVISER/MENTOR SUPPORT

Quality of mentoring from the Laboratory NRC Adviser (USMA Mentor, if applicable)

#### **Comments**

Dr Sciuto has provided me with excellent support in all aspects of laboratory work and research and career opportunities. I have been able to integrate my skill and mindset into an adaptive laboratory group and have been encouraged to contribute in any possible. He is the type of mentor that is able to help all personality types excel, and I have noted that in both myself and the people I work with in his group.

#### LPR SUPPORT

Quality of administrative support from the **L**aboratory (e.g., NIST, NRL, IWR, FHWA) NRC **P**rogram **R**epresentative (LPR)

Dr Kan has been available for all the necessary assistance I have needed in terms of communication between the institute and NRC.

#### NRC SUPPORT

Quality of administrative support. Please assess respective NRC aspects (e.g., moving company, insurance, Omega, payroll, coordinator, travel, etc.)

#### **Comments**

I was on a very tight schedule to move out when I received my appointment, and within 10 hours of having been confirmed, a everything concerning my move was sorted out. The moving company was excellent and reliable and the billing was easily sorted out with the NRC. The personnel in charge of communicating with me for health insurance were knowledgeable and of great assistance. Travel and payroll were also excellent in their communication and ability to rapidly process requests.

#### 18) PLEASE PROVIDE ANY SUGGESTIONS FOR PROGRAM IMPROVEMENT.

Communication on whether it was possible to obtain a security clearance through NRC was somewhat lacking, or uninforming. Clear delineations as to the policy would be helpful, especially to the institute.

Please do NOT scan to PDF. Send the Final Report as MSWord document via e-mail to your NRC Program Coordinator

No handwritten signature required; As but you may upload a scanned Lisignature file below: Ja

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